

FORM PTO-1390 (Modified)  
(REV 11-98)

U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE

TRANSMITTAL LETTER TO THE UNITED STATES  
DESIGNATED/ELECTED OFFICE (DO/EO/US)  
CONCERNING A FILING UNDER 35 U.S.C. 371

ATTORNEY'S DOCKET NUMBER

205554US0PCT

U.S. APPLICATION NO. (IF KNOWN, SEE 37 CFR

09/807962

INTERNATIONAL APPLICATION NO.  
PCT/EP99/08306INTERNATIONAL FILING DATE  
27 OCTOBER 1999PRIORITY DATE CLAIMED  
30 OCTOBER 1998

## TITLE OF INVENTION

2-AMINO-THIAZOLE DERIVATIVES, PROCESS FOR THEIR PREPARATION, AND THEIR USE AS  
ANTITUMOR AGENTSAPPLICANT(S) FOR DO/EO/US  
Paolo PEVARELLO, et al.

Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information:

1.  This is a **FIRST** submission of items concerning a filing under 35 U.S.C. 371.
2.  This is a **SECOND** or **SUBSEQUENT** submission of items concerning a filing under 35 U.S.C. 371.
3.  This is an express request to begin national examination procedures (35 U.S.C. 371(f)) at any time rather than delay examination until the expiration of the applicable time limit set in 35 U.S.C. 371(b) and PCT Articles 22 and 39(1).
4.  A proper Demand for International Preliminary Examination was made by the 19th month from the earliest claimed priority date.
5.  A copy of the International Application as filed (35 U.S.C. 371 (c) (2))
  - a.  is transmitted herewith (required only if not transmitted by the International Bureau).
  - b.  has been transmitted by the International Bureau.
  - c.  is not required, as the application was filed in the United States Receiving Office (RO/US).
6.  A translation of the International Application into English (35 U.S.C. 371(c)(2)).
7.  A copy of the International Search Report (PCT/ISA/210).
8.  Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371 (c)(3))
  - a.  are transmitted herewith (required only if not transmitted by the International Bureau).
  - b.  have been transmitted by the International Bureau.
  - c.  have not been made; however, the time limit for making such amendments has NOT expired.
  - d.  have not been made and will not be made.
9.  A translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371(c)(3)).
10.  An oath or declaration of the inventor(s) (35 U.S.C. 371 (c)(4)).
11.  A copy of the International Preliminary Examination Report (PCT/IEPA/409).
12.  A translation of the annexes to the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371 (c)(5)).

## Items 13 to 20 below concern document(s) or information included:

13.  An Information Disclosure Statement under 37 CFR 1.97 and 1.98.
14.  An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included.
15.  A **FIRST** preliminary amendment.
16.  A **SECOND** or **SUBSEQUENT** preliminary amendment.
17.  A substitute specification.
18.  A change of power of attorney and/or address letter.
19.  Certificate of Mailing by Express Mail
20.  Other items or information:

Request for Consideration of Documents Cited in International Search Report  
Notice of Priority  
PCT/IB/308

U.S. APPLICATION NO. (IF KNOWN) <b>72R 09/807962</b>		INTERNATIONAL APPLICATION NO. <b>PCT/EP99/08306</b>	ATTORNEY'S DOCKET NUMBER <b>205554US0PCT</b>																				
<p>21. The following fees are submitted:</p> <p><b>BASIC NATIONAL FEE (37 CFR 1.492 (a) (1) - (5)) :</b></p> <table border="0"> <tr> <td><input type="checkbox"/> Neither international preliminary examination fee (37 CFR 1.482) nor international search fee (37 CFR 1.445(a)(2) paid to USPTO and International Search Report not prepared by the EPO or JPO .....</td> <td style="text-align: right;"><b>\$1,000.00</b></td> </tr> <tr> <td><input checked="" type="checkbox"/> International preliminary examination fee (37 CFR 1.482) not paid to USPTO but International Search Report prepared by the EPO or JPO .....</td> <td style="text-align: right;"><b>\$860.00</b></td> </tr> <tr> <td><input type="checkbox"/> International preliminary examination fee (37 CFR 1.482) not paid to USPTO but international search fee (37 CFR 1.445(a)(2)) paid to USPTO .....</td> <td style="text-align: right;"><b>\$710.00</b></td> </tr> <tr> <td><input type="checkbox"/> International preliminary examination fee paid to USPTO (37 CFR 1.482) but all claims did not satisfy provisions of PCT Article 33(1)-(4) .....</td> <td style="text-align: right;"><b>\$690.00</b></td> </tr> <tr> <td><input type="checkbox"/> International preliminary examination fee paid to USPTO (37 CFR 1.482) and all claims satisfied provisions of PCT Article 33(1)-(4) .....</td> <td style="text-align: right;"><b>\$100.00</b></td> </tr> </table> <p><b>ENTER APPROPRIATE BASIC FEE AMOUNT =</b> <input type="text" value="860.00"/></p> <p>Surcharge of <b>\$130.00</b> for furnishing the oath or declaration later than <input type="checkbox"/> 20 <input type="checkbox"/> 30 months from the earliest claimed priority date (37 CFR 1.492 (e)). <input type="text" value="0.00"/></p>				<input type="checkbox"/> Neither international preliminary examination fee (37 CFR 1.482) nor international search fee (37 CFR 1.445(a)(2) paid to USPTO and International Search Report not prepared by the EPO or JPO .....	<b>\$1,000.00</b>	<input checked="" type="checkbox"/> International preliminary examination fee (37 CFR 1.482) not paid to USPTO but International Search Report prepared by the EPO or JPO .....	<b>\$860.00</b>	<input type="checkbox"/> International preliminary examination fee (37 CFR 1.482) not paid to USPTO but international search fee (37 CFR 1.445(a)(2)) paid to USPTO .....	<b>\$710.00</b>	<input type="checkbox"/> International preliminary examination fee paid to USPTO (37 CFR 1.482) but all claims did not satisfy provisions of PCT Article 33(1)-(4) .....	<b>\$690.00</b>	<input type="checkbox"/> International preliminary examination fee paid to USPTO (37 CFR 1.482) and all claims satisfied provisions of PCT Article 33(1)-(4) .....	<b>\$100.00</b>										
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<p><input checked="" type="checkbox"/> A check in the amount of <b>\$860.00</b> to cover the above fees is enclosed.</p> <p><input type="checkbox"/> Please charge my Deposit Account No. <b>15-0030</b> in the amount of <b>\$</b> to cover the above fees. A duplicate copy of this sheet is enclosed.</p> <p><input type="checkbox"/> The Commissioner is hereby authorized to charge any fees which may be required, or credit any overpayment to Deposit Account No. <b>15-0030</b> A duplicate copy of this sheet is enclosed.</p>																							
<p>NOTE: Where an appropriate time limit under 37 CFR 1.494 or 1.495 has not been met, a petition to revive (37 CFR 1.137(a) or (b)) must be filed and granted to restore the application to pending status.</p>																							
<p>SEND ALL CORRESPONDENCE TO:</p> <div style="border: 1px solid black; padding: 10px; text-align: center;">   <b>22850</b> </div>																							
<div style="text-align: right; margin-right: 100px;">   <b>SIGNATURE</b>  <b>Norman F. Oblon</b>  <b>NAME</b>  <b>24,618</b>  <b>REGISTRATION NUMBER</b>  <b>Apri 26 2001</b>  <b>DATE</b> </div>																							

IN THE UNITED STATES PATENT & TRADEMARK OFFICEIN RE APPLICATION OF :  
PAOLO PEVARELLO ET AL

: ATTN: APPLICATION DIVISION

SERIAL NO: NEW US PCT APPLN.  
(Based on PCT NO:EP99/08306)

:

FILED: HEREWITH : EXAMINER:

FOR: 2-AMINO-THIAZOLE DERIVATIVES,  
PROCESS FOR THEIR PREPARATION,  
AND THEIR USE AS ANTITUMOR AGENTSPRELIMINARY AMENDMENTASSISTANT COMMISSIONER FOR PATENTS  
WASHINGTON, D.C. 20231

SIR:

Prior to examination on the merits, please amend the above-identified application as follows:

IN THE CLAIMS

Please amend the claims as shown in the marked-up copy to read as follows:

--5. (Amended) Use according to claim 1 wherein the medicament enables tumor angiogenesis and metastasis inhibition.

11. (Amended) A compound of formula (I) according to claim 6, whenever appropriate in the form of a pharmaceutically acceptable salt, selected from the group consisting of:

1. ethyl 3-[(5-bromo-1,3-thiazol-2-yl)amino]-3-oxopropanoate;
2. N-(5-bromo-1,3-thiazol-2-yl)-2-phenyl-acetamide;

3. N-(5-bromo-1,3-thiazol-2-yl)-benzamide;  
4. Ethyl 4-[(5-bromo-1,3-thiazol-2-yl)amino]-4-oxobutanoate;  
5. N-(5-Bromo-thiazol-2-yl)-3-hydroxy-propionamide;  
6. N-(5-Bromo-1,3-thiazol-2-yl)-4-hydroxybutanamide;  
7. N-(5-Bromo-thiazol-2-yl)-2-ethoxy-acetamide;  
8. 2-N-[2-(3-pyridyl)-acetyl-amino]-5-bromo-thiazole;  
9. 2-N-[2-(3-pyridyl)-acetyl-amino]-5-isopropyl-thiazole;  
10.N-(5-bromo-1,3-thiazol-2-yl)-2-(3-hydroxyphenyl)acetamide;  
11.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-hydroxyphenyl)acetamide;  
12.N-(5-bromo-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide;  
13.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide;  
14.N-(5-bromo-1,3-thiazol-2-yl)-2-(3-chlorophenyl)acetamide;  
15.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-chlorophenyl)acetamide;  
16.N-(5-bromo-1,3-thiazol-2-yl)-2-(4-hydroxyphenyl)acetamide;  
17.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxyphenyl)acetamide;  
18.N-(5-bromo-1,3-thiazol-2-yl)-2-(3,4-dihydroxyphenyl)acetamide;  
19.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dihydroxyphenyl)acetamide;  
20.N-(5-bromo-1,3-thiazol-2-yl)-2-(4-hydroxy-3-methoxyphenyl)acetamide;  
21.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxy-3-methoxyphenyl)acetamide;  
22.N-(5-bromo-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide;  
23.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide;  
24.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-chlorophenyl)acetamide;  
25.N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-acetamide;  
26.N-(5-bromo-thiazol-2-yl)-4-sulfamoyl-benzamide;

27.N-(5-isopropyl-thiazol-2-yl)-4-sulfamoyl-benzamide;  
28.4-amino-N-(5-bromo-1,3-thiazol-2-yl)butanamide;  
29.3-amino-N-(5-bromo-1,3-thiazol-2-yl)propionamide;  
30.N-(5-isopropyl-1,3-thiazol-2-yl)-butanamide;  
31.N-(5-bromo-1,3-thiazol-2-yl)-butanamide;  
32.N-(5-chloro-1,3-thiazol-2-yl)-butanamide;  
33.N-(5-phenyl-1,3-thiazol-2-yl)-butanamide;  
34.N-(5-nitro-1,3-thiazol-2-yl)-butanamide;  
35.N-(5-methyl-1,3-thiazol-2-yl)-butanamide;  
36.N-(5-benzyl-1,3-thiazol-2-yl)-butanamide;  
37.N-(5-isobutyl-1,3-thiazol-2-yl)-butanamide;  
38.N-(5-cyclopropyl-1,3-thiazol-2-yl)-butanamide;  
39.N-[5-[2-(methylsulfonyl)ethyl]-1,3-thiazol-2-yl]-butanamide;  
40.N-[5-(2-methylthioethyl)-1,3-thiazol-2-yl]-butanamide;  
41.N-[5-[2-(methoxycarbonyl)ethyl]-1,3-thiazol-2-yl]-butanamide;  
42.N-[5-(3-methoxy-propyl)-1,3-thiazol-2-yl]-butanamide;  
43.N-[5-(2-ethoxy-ethyl)-1,3-thiazol-2-yl]-butanamide;  
44.N-[5-(indol-3-yl-methyl)-1,3-thiazol-2-yl]-butanamide;  
45.N-[5-(3-oxo-butyl)-1,3-thiazol-2-yl]-butanamide;  
46.2-[3-(3-chloropropoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;  
47.2-[3-(2-chloroethoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;  
48.2-(4-aminophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;  
49.4-amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
50.2-(2-amino-1,3-thiazol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;

51.N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[3-(4-morpholinyl)propoxy]phenyl}acetamide;  
52.N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[2-(4-morpholinyl)ethoxy]phenyl}acetamide;  
53.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-[3-(1-pirrolidinyl)propoxy]phenyl)acetamide;  
54.N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[3-(4-methyl-lpiperazinyl)propoxy]phenyl}acetamide;  
55.2-{3-[2-(dimethylamino)ethoxy]phenyl}-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;  
56.2-{3-[3-(dimethylamino)propoxy]phenyl}-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
57.2-[4-(dimethylamino)phenyl]-N-(5-isobutyl-1,3-thiazol-2-yl)acetamide  
58.2-(1,3-benzodioxol-5-yl)-N-(5-isobutyl-1,3-thiazol-2-yl)acetamide  
59.N-(5-benzyl-1,3-thiazol-2-yl)-2-[4-(dimethylamino)phenyl]acetamide  
60.N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(2-methoxyethoxy)phenyl]acetamide  
61.3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-methyl-lpiperazinyl)benzamide  
62.N-(5-isobutyl-1,3-thiazol-2-yl)-2-(3-pyridinyl)acetamide  
63.N-(5-benzyl-1,3-thiazol-2-yl)-2-(3-pyridinyl)acetamide  
64.2-[N-[2'-N'-(ethoxycarbonyl-methyl)-amino]-acetyl]amino-5-bromo-thiazole  
65.2-anilino-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
66.(R)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide  
67.(S)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide  
68.N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
69.2,5-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
70.3,5-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
71.3,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
72.2,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
73.2,3-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

74.3-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
75.2-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
76.4-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
77.3-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
78.4-chloro-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
79.5-bromo-2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
80.3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
81.2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
82.4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
83.3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
84.2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
85.4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
86.2,4-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
87.3,4-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
88.2,3,4,5,6-pentafluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
89.N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-3-nitrobenzamide  
90.N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-nitrobenzamide  
91.N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-nitrobenzamide  
92.N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-nitrobenzamide  
93.N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxy-2-nitrobenzamide  
94.N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-2-nitrobenzamide  
95.N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxy-3-nitrobenzamide  
96.N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-nitrobenzamide  
97.N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dinitrobenzamide

98.5-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-2-nitrophenyl octanoate

99.N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide

100. N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide

101. N-(5-isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamide

102. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl)-3-nitrobenzamide

103. 4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide

104. 6-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide

105. 4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide

106. 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamide

107. 5-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide

108. 2-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-5-nitrobenzamide

109. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide

110. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide

111. N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitro-4-(trifluoromethyl)benzamide

112. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-bis(trifluoromethyl)benzamide

113. N-(5-isopropyl-1,3-thiazol-2-yl)-2,6-bis(trifluoromethyl)benzamide

114. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(trifluoromethyl)benzamide

115. N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide

116. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethyl)benzamide

117. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide

118. 5-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide

119. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethyl)benzamide

120. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide

121. methyl 4-((5-isopropyl-1,3-thiazol-2-yl)amino)carbonyl}benzoate

122. methyl 2-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)benzoate  
123. 4-cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
124. 3-cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
125. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methylbenzamide  
126. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methylbenzamide  
127. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylbenzamide  
128. N-(5-isopropyl-1,3-thiazol-2-yl)-4-vinylbenzamide  
129. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(2-phenylethynyl)benzamide  
130. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-methylbenzamide  
131. 2-benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
132. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenethylbenzamide  
133. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylbenzamide  
134. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenylbenzamide  
135. 4-(tert-butyl)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
136. N-(5-isopropyl-1,3-thiazol-2-yl)-4-isopropylbenzamide  
137. N-(5-isopropyl-1,3-thiazol-2-yl)-4-pentylbenzamide  
138. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylbenzamide  
139. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4-dimethylbenzamide  
140. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethylbenzamide  
141. 4-acetyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
142. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl)benzamide  
143. 5-(aminosulfonyl)-2,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
144. 5-(aminosulfonyl)-4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
145. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide

146. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide  
147. 5-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxybenzamide  
148. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide  
149. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxybenzamide  
150. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxybenzamide  
151. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4-dimethoxybenzamide  
152. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethoxybenzamide  
153. N-(5-isopropyl-1,3-thiazol-2-yl)-2,4-dimethoxybenzamide  
154. N-(5-isopropyl-1,3-thiazol-2-yl)-2,3-dimethoxybenzamide  
155. N-(5-isopropyl-1,3-thiazol-2-yl)-3-phenoxybenzamide  
156. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenoxybenzamide  
157. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenoxybenzamide  
158. 2-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
159. 4-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
160. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4,5-trimethoxybenzamide  
161. 3,4-diethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
162. 3,4,5-triethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
163. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4(methoxymethoxy)benzamide  
164. 4-butoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
165. N-(5-isopropyl-1,3-thiazol-2-yl)-4-propoxybenzamide  
166. 4-isopropoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
167. N-(5-isopropyl-1,3-thiazol-2-yl)-1,3-benzodioxole-5-carboxamide  
168. 4-(benzyloxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
169. 4-(2-cyclohexen-1-ylloxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

170. N-(5-isopropyl-1,3-thiazol-2-yl)-4(trifluoromethoxy)benzamide  
171. 4-(difluoromethoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
172. N-(5-isopropyl-1,3-thiazol-2-yl)-4(methylsulfanyl)benzamide  
173. 2-[(4-chlorophenyl)sulfinyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
174. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[(4-nitrophenyl)sulfinyl]benzamide  
175. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(4-methylphenyl)sulfonyl]-3-nitrobenzamide  
176. N-(5-isopropyl-1,3-thiazol-2-yl)-3-[(trifluoromethyl)sulfanyl]benzamide  
177. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxy-4-(methylsulfanyl)benzamide  
178. 2-[(2-cyanophenyl)sulfanyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
179. N~1~,N-1~-diethyl-3,6-difluoro-N-2~-(5-isopropyl-1,3-thiazol-2-yl)phthalamide  
180. 4-formyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
181. 2-formyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
182. 4-{[(2,5-dimethoxyanilino)carbonyl]amino}-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
183. 4(hydroxymethyl)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
184. 4-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-2-nitrobenzyl acetate  
185. 4-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-2-nitrobenzyl 4-(acetylamino)-3-iodobenzoate  
186. 4-(acetylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
187. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(2-phenylacetyl)amino]benzamide  
188. 4-(acetylamino)-3-iodo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
189. 4-amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
190. 4-(dimethylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
191. 3-(dimethylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
192. 2-(methylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

193. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(trifluoromethyl)anilino]benzamide

194. 3-{{[5-bromo-1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)methyl]amino}-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

195. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(1H-pyrrol-1-yl)benzamide

196. 2,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)isonicotinamide

197. 2-(4-bromophenyl)-6-(4-iodophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)isonicotinamide.

198. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(trifluoromethyl)anilino]nicotinamide

199. 2,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide

200. 5,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide

201. 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-6-methylnicotinamide

202. 2,6-dichloro-5-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide

203. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenoxy nicotinamide

204. N-(5-isopropyl-1,3-thiazol-2-yl)-6-(2,2,2-trifluoroethoxy)nicotinamide

205. N-(5-isopropyl-1,3-thiazol-2-yl)-2,6-dimethoxy nicotinamide

206. N-(5-isopropyl-1,3-thiazol-2-yl)-2-quinoxalinecarboxamide

207. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-pyrazinecarboxamide

208. N-(5-isopropyl-1,3-thiazol-2-yl)-8-quinolinecarboxamide

209. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-4-quinolinecarboxamide

210. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-1-phenyl-1H-pyrazole-4-carboxamide

211. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-1H-pyrazole-3-carboxamide

212. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-pyrazole-4-carboxamide

213. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-phenyl-2H-1,2,3-triazole-4-carboxamide

214. 2-[2-(2,1,3-benzoxadiazol-5-yloxy)methyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-1,3-thiazole-5-carboxamide

215. N-(5-isopropyl-1,3-thiazol-2-yl)-9H-fluorene-1-carboxamide

216. N-(5-isopropyl-1,3-thiazol-2-yl)-7-methoxy-1-benzofuran-2-carboxamide

217. N-(5-isopropyl-1,3-thiazol-2-yl)-1-[(4-methylphenyl)sulfonyl]-1H-pyrrole-3-carboxamide

218. 2-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamide

219. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamide

220. N-(5-isopropyl-1,3-thiazol-2-yl)-2-naphthamide

221. N-(5-isopropyl-1,3-thiazol-2-yl)-9,10-dioxo-9,10-dihydro-2-anthracenecarboxamide

222. N-(5-isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluorene-4-carboxamide

223. N-(5-isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluorene-1-carboxamide

224. N-(5-isopropyl-1,3-thiazol-2-yl)-8-oxo-5,6,7,8-tetrahydro-2-naphthalenecarboxamide

225. N-(5-isopropyl-1,3-thiazol-2-yl)-1,3-dioxo-1,3-dihydro-2-benzofuran-5-carboxamide

226. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-5-carboxamide

227. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-4-carboxamide

228. N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-2-phenyl-1H-indole-5-carboxamide

229. 2-butyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indole-5-carboxamide

230. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-6-carboxamide

231. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methoxy-1H-indole-2-carboxamide

232. 1-allyl-2-butyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-5-carboxamide

233. N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indole-2-carboxamide

234. 1-benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-1H-indole-5-carboxamide

235. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-1,2,3-benzotriazole-5-carboxamide

236. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-isoxazolecarboxamide

237. N-(5-isopropyl-1,3-thiazol-2-yl)-3-thiophenecarboxamide

238. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-thiophenecarboxamide  
239. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-thiophenecarboxamide  
240. 5-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-2-thiophenecarboxamide  
241. N-(5-isopropyl-1,3-thiazol-2-yl)-3-[(2,3,3-trichloroacryloyl)amino]-2-thiophenecarboxamide  
242. 5-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide  
243. N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide  
244. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(4-nitrophenyl)-2-furamide  
245. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(2-nitrophenyl)-2-furamide  
246. 5-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide  
247. N-(5-isopropyl-1,3-thiazol-2-yl)-5-[3-(trifluoromethyl)phenyl]-2-furamide  
248. 5-(4-chloro-2-nitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide  
249. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(4-methyl-2-nitrophenyl)-2-furamide  
250. 5-[2-chloro-5-(trifluoromethyl)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide  
251. *tert*-butyl (1*R*)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethylcarbamate  
252. (1*R*)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate  
253. (1*S*)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate  
254. (R,S)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide  
255. (R)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide  
256. (S)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide  
257. 2-(acetylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide  
258. (R,S)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide  
259. (R)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide  
260. (S)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

261. 3,3,3-trifluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxy-2-phenylpropanamide  
262. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(1-naphthyl)acetamide  
263. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-naphthyl)acetamide  
264. 2-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
265. 2-(1,3-benzodioxol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
266. 2-(2,4-dinitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
267. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-methyl-1H-indol-3-yl)acetamide  
268. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(1-methyl-1H-indol-3-yl)acetamide  
269. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(5-methoxy-1H-indol-3-yl)acetamide  
270. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(5-benzyloxy-1H-indol-3-yl)acetamide  
271. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxy-2-methyl-1H-indol-3-yl)acetamide  
272. 2-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxoacetamide  
273. 2-(5-bromo-1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
274. 2-(5-fluoro-1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
275. 2-[1-(4-chlorobenzoyl)-5-methoxy-2-methyl-1H-indol-3-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
276. 3-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide  
277. 4-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)butanamide  
278. N-(5-isopropyl-1,3-thiazol-2-yl)-3-(2-thienyl)propanamide  
279. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-thienyl)acetamide  
280. N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxo-2-(2-thienyl)acetamide  
281. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-thienyl)acetamide  
282. 2-(5-chloro-1-benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
283. 2-(1-benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

284. 2-[2-(formylamino)-1,3-thiazol-4-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamide

285. 2-{2-[(2-chloroacetyl)amino]-1,3-thiazol-4-yl}-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamide

286. 2-chloro-N-(4-(2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl)-1,3-thiazol-2-yl)acetamide

287. ethyl 2-({[2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-(1H-pyrazol-3-yl)ethylidene]amino}oxy)acetate

288. 2-(2-furyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxoacetamide

289. 2-(5-bromo-3-pyridinyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

290. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(7-methoxy-2-oxo-2H-chromen-4-yl)acetamide

291. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenyl-3-butenamide

292. N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxo-4-(4-methyl-phenyl)butanamide

293. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-nitrophenyl)butanamide

294. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenylbutanamide

295. benzyl 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]-4-oxobutylcarbamate

296. methyl 5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate

297. 4-(1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)butanamide

298. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-methoxy-1-naphthyl)-4-oxobutanamide

299. 3-(2-chlorophenoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

300. 3-(4-methylphenoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

301. 3-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

302. 3-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

303. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylpentanamide

304. 3-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide  
305. 3-(4-methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide  
306. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide  
307. 3-phenyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide  
308. 2-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
309. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methylbutanamide  
310. N-(5-isopropyl-1,3-thiazol-2-yl)-5-oxo-5-phenylpentanamide  
311. 2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl phenyl acetate  
312. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[4-(1-oxo-1,3-dihydro-2H-isoindol-2-yl)phenyl]propanamide  
313. 1-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)cyclopentanecarboxamide  
314. 1-phenyl-N-(5-isopropyl-1,3-thiazol-2-yl)cyclopentanecarboxamide  
315. 2-(3-bromo-4-methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
316. 2-(2-nitro-4-trifluoromethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
317. 5-cyclohexyl 1-(4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}benzyl) (2S)-2-[(tert-butoxycarbonyl)amino]pentanedioate  
318. 2-(5,6-dimethyl-1H-benzimidazol-1-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
319. 2-[5-(4-chlorophenyl)-2H-1,2,3,4-tetraazol-2-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
320. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[5-(1-pyrrolidinyl)-2H-1,2,3,4-tetraazol-2-yl]acetamide  
321. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methyl-1-benzothiophen-2-yl)acetamide  
322. N-(5-isopropyl-1,3-thiazol-2-yl)-4,4-bis(4-methylphenyl)-3-butenamide  
323. 2-cyclopropyl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

324. N-{4-bromo-6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexyl}benzamide  
325. 2-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
326. benzyl 6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexylcarbamate  
327. N-1- (5-isopropyl-1,3-thiazol-2-yl)-N-4- (2-propynyl)-2-butenediamide  
328. 4-(2,4-dimethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide  
329. 4-(4-benzyloxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide  
330. 4-(thiphen-2-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide  
331. benzyl 2-{[(benzyloxy)carbonyl]amino}-5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate  
332. 4-(1H-indol-3-yl)-N-{3-[(5-isopropyl-1,3-thiazol-2-yl)amino]-3-oxopropyl}butanamide  
333. 4-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}phenyl 4-chlorobenzenesulfonate  
334. N-(5-isopropyl-1,3-thiazol-2-yl)-4-{[(2-methoxyanilino)carbonyl]amino}benzamide  
335. 4-{[2-(isopropylsulfonyl)acetyl]amino}-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
336. N-(5-isopropyl-1,3-thiazol-2-yl)-4-{[2-(phenylsulfanyl)acetyl]amino}benzamide  
337. 4-[(diethylamino)sulfonyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
338. 2-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
339. 3,5-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
340. 3-[(2-fluoroanilino)carbonyl]amino}-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
341. N-(5-isopropyl-1,3-thiazol-2-yl)-1-phenyl-5-propyl-1H-pyrazole-4-carboxamide  
342. 3-chloro-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfanyl)-2-thiophenecarboxamide  
343. 3-iodo-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfanyl)-2-thiophenecarboxamide

344. 2-[(4-chlorophenyl)sulfonyl]methyl}-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-1,3-thiazole-5-carboxamide

345. 5-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(trifluoromethyl)-3-furamide

346. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,3,4,5,6-pentafluorophenyl)acetamide

347. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-fluorophenyl)acetamide

348. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-bromophenyl)acetamide

349. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-chlorophenyl)acetamide

350. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-nitrophenyl) acetamide

351. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-trifluoromethylphenyl)acetamide

352. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-methoxyphenyl)acetamide

353. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-dimethoxyphenyl)acetamide

354. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-difluorophenyl)acetamide

355. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4,5-trimethoxyphenyl)acetamide

356. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,6-dichlorophenyl)acetamide

357. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-chloro-6-fluorophenyl)acetamide

358. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,5-dimethoxyphenyl)acetamide

359. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,5-difluorophenyl)acetamide

360. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-bistrifluoromethylphenyl)acetamide

361. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methylthiophenyl)acetamide

362. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide

363. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-bromophenyl)acetamide

364. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-chlorophenyl)acetamide

365. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-fluorophenyl)acetamide

366. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl)acetamide

367. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-trifluoromethylphenyl)acetamide  
368. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methylphenyl)acetamide  
369. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-dimethylaminophenyl)acetamide  
370. 2-[1,1'-biphenyl]-4-yl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide,  
371. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-trifluoromethylphenyl)acetamide  
372. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-bromophenyl)acetamide  
373. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-chlorophenyl)acetamide  
374. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-nitrophenyl)acetamide  
375. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide  
376. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-dinitrophenyl)acetamide  
377. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-dichlorophenyl)acetamide  
378. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-difluorophenyl)acetamide  
379. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-benzyloxy-3-methoxyphenyl)acetamide  
380. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dichlorophenyl)acetamide  
381. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-difluorophenyl)acetamide  
382. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dimethoxyphenyl)acetamide  
383. 2-(2,3-dihydro-1H-inden-5-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
384. N-(5-isopropyl-1,3-thiazol-2-yl)-1-phenylcyclopropanecarboxamide  
385. 2-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide  
386. 2-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide  
387. N-(5-isopropyl-1,3-thiazol-2-yl)-2,2-diphenylacetamide  
388. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-nitrophenoxy)acetamide  
389. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl)propanamide  
390. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide

391. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-isobutylphenyl)propanamide

392. N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxo-2-phenylacetamide

393. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-phenylpentanamide

394. (E, Z)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-2-butenamide

395. N-(5-isopropyl-1,3-thiazol-2-yl)bicyclo[4.2.0]octa-1,3,5-triene-7-carboxamide

396. N-(5-isopropyl-1,3-thiazol-2-yl)-3-oxo-1-indanecarboxamide

397. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl)butanamide

398. tert-butyl (1S)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-1-methyl-2-oxoethylcarbamate

399. tert-butyl (1S,2S)-1-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-2-methylbutylcarbamate

400. tert-butyl 2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

401. tert-butyl (1S)-5-amino-1-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}pentylcarbamate

402. tert-butyl 4-[(imino {[(4-methylphenyl)sulfonyl]amino}methyl)amino]-1-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}butylcarbamate

403. tert-butyl 1-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-3-(tritylamino)propylcarbamate

404. tert-butyl (1S)-1-(benzyloxymethyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

405. tert-butyl (1S)-1-benzyl-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

406. tert-butyl (1R)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-(benzylthiomethyl)ethylcarbamate

407. benzyl (3S)-3-[(tert-butoxycarbonyl)amino]-4-[(5-isopropyl-1,3-thiazol-2-yl)amino]-4-oxobutanoate

408. *tert*-butyl (2*S*)-2-[(5-isopropyl-1,3-thiazol-2-yl)aminolcarbonyl]-1-pyrrolidinecarboxylate

409. *tert*-butyl (1*S*)-1-(1*H*-indol-3-ylmethyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

410. *tert*-butyl (1*S*)-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)-3-(methylsulfanyl)propylcarbamate

411. *tert*-butyl (1*S*)-2-benzyloxy-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)propylcarbamate

412. *tert*-butyl (1*S*)-1-(4-benzyloxybenzyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

413. *tert*-butyl (1*S*)-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)-2-methylpropylcarbamate

414. *tert*-butyl (1*S*)-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)-3-methylbutylcarbamate

415. benzyl (4*S*)-4-[(*tert*-butoxycarbonyl)amino]-5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate

and the pharmaceutically acceptable salts thereof.--

REMARKS

Claims 1-14 are active in the present application. The claims are amended to remove multiple dependencies. No new matter is added. An action on the merits and allowance of the claims is solicited.

Respectfully submitted,

OBLON, SPIVAK, McCLELLAND,  
MAIER & NEUSTADT, P.C.



Norman F. Oblon  
Attorney of Record  
Registration No. 24,618

Daniel J. Pereira  
Registration No. 45,518



**22850**

(703) 413-3000  
Fax #: (703)413-2220  
DJPER/rac  
H:\205554US-PR.wpd

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Amendment Filed on: \_\_\_\_\_  
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IN THE CLAIMS

Please amend the claims as follows:

--5. (Amended) Use according to [any one of the preceding claims] claim 1 wherein the medicament enables tumor angiogenesis and metastasis inhibition.

11. (Amended) A compound of formula (I) according to [any one of the preceding claims] claim 6, whenever appropriate in the form of a pharmaceutically acceptable salt, selected from the group consisting of:

1. ethyl 3-[(5-bromo-1,3-thiazol-2-yl)amino]-3-oxopropanoate;
2. N-(5-bromo-1,3-thiazol-2-yl)-2-phenyl-acetamide;
3. N-(5-bromo-1,3-thiazol-2-yl)-benzamide;
4. Ethyl 4-[(5-bromo-1,3-thiazol-2-yl)amino]-4-oxobutanoate;
5. N-(5-Bromo-thiazol-2-yl)-3-hydroxy-propionamide;
6. N-(5-Bromo-1,3-thiazol-2-yl)-4-hydroxybutanamide;
7. N-(5-Bromo-thiazol-2-yl)-2-ethoxy-acetamide;
8. 2-N-[2-(3-pyridyl)-acetyl-amino]-5-bromo-thiazole;
9. 2-N-[2-(3-pyridyl)-acetyl-amino]-5-isopropyl-thiazole;
- 10.N-(5-bromo-1,3-thiazol-2-yl)-2-(3-hydroxyphenyl)acetamide;
- 11.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-hydroxyphenyl)acetamide;
- 12.N-(5-bromo-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide;

13.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide;

14.N-(5-bromo-1,3-thiazol-2-yl)-2-(3-chlorophenyl)acetamide;

15.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-chlorophenyl)acetamide;

16.N-(5-bromo-1,3-thiazol-2-yl)-2-(4-hydroxyphenyl)acetamide;

17.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxyphenyl)acetamide;

18.N-(5-bromo-1,3-thiazol-2-yl)-2-(3,4-dihydroxyphenyl)acetamide;

19.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dihydroxyphenyl)acetamide;

20.N-(5-bromo-1,3-thiazol-2-yl)-2-(4-hydroxy-3-methoxyphenyl)acetamide;

21.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxy-3-methoxyphenyl)acetamide;

22.N-(5-bromo-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide;

23.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide;

24.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-chlorophenyl)acetamide;

25.N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-acetamide;

26.N-(5-bromo-thiazol-2-yl)-4-sulfamoyl-benzamide;

27.N-(5-isopropyl-thiazol-2-yl)-4-sulfamoyl-benzamide;

28.4-amino-N-(5-bromo-1,3-thiazol-2-yl)butanamide;

29.3-amino-N-(5-bromo-1,3-thiazol-2-yl)propionamide;

30.N-(5-isopropyl-1,3-thiazol-2-yl)-butanamide;

31.N-(5-bromo-1,3-thiazol-2-yl)-butanamide;

32.N-(5-chloro-1,3-thiazol-2-yl)-butanamide;

33.N-(5-phenyl-1,3-thiazol-2-yl)-butanamide;

34.N-(5-nitro-1,3-thiazol-2-yl)-butanamide;

35.N-(5-methyl-1,3-thiazol-2-yl)-butanamide;

36.N-(5-benzyl-1,3-thiazol-2-yl)-butanamide;

37.N-(5-isobutyl-1,3-thiazol-2-yl)-butanamide;  
38.N-(5-cyclopropyl-1,3-thiazol-2-yl)-butanamide;  
39.N-{5-[2-(methylsulfonyl)ethyl]-1,3-thiazol-2-yl}-butanamide;  
40.N-[5-(2-methylthioethyl)-1,3-thiazol-2-yl]-butanamide;  
41.N-{5-[2-(methoxycarbonyl)ethyl]-1,3-thiazol-2-yl}-butanamide;  
42.N-[5-(3-methoxy-propyl)-1,3-thiazol-2-yl]-butanamide;  
43.N-[5-(2-ethoxy-ethyl)-1,3-thiazol-2-yl]-butanamide;  
44.N-[5-(indol-3-yl-methyl)-1,3-thiazol-2-yl]-butanamide;  
45.N-[5-(3-oxo-butyl)-1,3-thiazol-2 yl]-butanamide;  
46.2-[3-(3-chloropropoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;  
47.2-[3-(2-chloroethoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;  
48.2-(4-aminophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;  
49.4-amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
50.2-(2-amino-1,3-thiazol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;  
51.N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[3-(4-morpholinyl)propoxy]phenyl}acetamide;  
52.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-[2-(4-morpholinyl)ethoxy]phenyl)acetamide;  
53.N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-[3-(1-pirrolidinyl)propoxy]phenyl)acetamide;  
54.N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[3-(4-methyl-lpiperazinyl)propoxy]phenyl}acetamide;  
55.2-{3-[2-(dimethylamino)ethoxy]phenyl}-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;  
56.2-{3-[3-(dimethylamino)propoxy]phenyl}-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
57.2-[4-(dimethylamino)phenyl]-N-(5-isobutyl-1,3-thiazol-2-yl)acetamide  
58.2-(1,3-benzodioxol-5-yl)-N-(5-isobutyl-1,3-thiazol-2-yl)acetamide  
59.N-(5-benzyl-1,3-thiazol-2-yl)-2-[4-(dimethylamino)phenyl]acetamide

60.N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(2-methoxyethoxy)phenyl]acetamide  
61.3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-methyl-1-piperazinyl)benzamide  
62.N-(5-isobutyl-1,3-thiazol-2-yl)-2-(3-pyridinyl)acetamide  
63.N-(5-benzyl-1,3-thiazol-2-yl)-2-(3-pyridinyl)acetamide  
64.2-[N-[2'-N'-(ethoxycarbonyl-methyl)-amino]-acetyl]amino-5-bromo-thiazole  
65.2-anilino-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
66.(R)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide  
67.(S)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide  
68.N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
69.2,5-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
70.3,5-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
71.3,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
72.2,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
73.2,3-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
74.3-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
75.2-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
76.4-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
77.3-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
78.4-chloro-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
79.5-bromo-2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
80.3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
81.2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
82.4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
83.3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

84. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
85. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
86. 2,4-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
87. 3,4-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
88. 2,3,4,5,6-pentafluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
89. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-3-nitrobenzamide  
90. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-nitrobenzamide  
91. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-nitrobenzamide  
92. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-nitrobenzamide  
93. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxy-2-nitrobenzamide  
94. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-2-nitrobenzamide  
95. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxy-3-nitrobenzamide  
96. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-nitrobenzamide  
97. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dinitrobenzamide  
98. 5-{{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-2-nitrophenyl} octanoate  
99. N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide  
100. N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide  
101. N-(5-isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamide  
102. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl)-3-nitrobenzamide  
103. 4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide  
104. 6-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide  
105. 4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide  
106. 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamide  
107. 5-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide

108. 2-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-5-nitrobenzamide  
109. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide  
110. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide  
111. N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitro-4-(trifluoromethyl)benzamide  
112. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-bis(trifluoromethyl)benzamide  
113. N-(5-isopropyl-1,3-thiazol-2-yl)-2,6-bis(trifluoromethyl)benzamide  
114. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(trifluoromethyl)benzamide  
115. N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide  
116. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethyl)benzamide  
117. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide  
118. 5-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide  
119. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethyl)benzamide  
120. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide  
121. methyl 4-([(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)benzoate  
122. methyl 2-([(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)benzoate  
123. 4-cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
124. 3-cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
125. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methylbenzamide  
126. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methylbenzamide  
127. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylbenzamide  
128. N-(5-isopropyl-1,3-thiazol-2-yl)-4-vinylbenzamide  
129. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(2-phenylethynyl)benzamide  
130. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-methylbenzamide  
131. 2-benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

132. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenethylbenzamide  
133. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylbenzamide  
134. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenylbenzamide  
135. 4-(tert-butyl)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
136. N-(5-isopropyl-1,3-thiazol-2-yl)-4-isopropylbenzamide  
137. N-(5-isopropyl-1,3-thiazol-2-yl)-4-pentylbenzamide  
138. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylbenzamide  
139. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4-dimethylbenzamide  
140. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethylbenzamide  
141. 4-acetyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
142. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl)benzamide  
143. 5-(aminosulfonyl)-2,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
144. 5-(aminosulfonyl)-4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
145. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide  
146. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide  
147. 5-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxybenzamide  
148. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide  
149. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxybenzamide  
150. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxybenzamide  
151. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4-dimethoxybenzamide  
152. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethoxybenzamide  
153. N-(5-isopropyl-1,3-thiazol-2-yl)-2,4-dimethoxybenzamide  
154. N-(5-isopropyl-1,3-thiazol-2-yl)-2,3-dimethoxybenzamide  
155. N-(5-isopropyl-1,3-thiazol-2-yl)-3-phenoxybenzamide

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156. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenoxybenzamide
157. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenoxybenzamide
158. 2-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
159. 4-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
160. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4,5-trimethoxybenzamide
161. 3,4-diethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
162. 3,4,5-triethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
163. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4(methoxymethoxy)benzamide
164. 4-butoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
165. N-(5-isopropyl-1,3-thiazol-2-yl)-4-propoxybenzamide
166. 4-isopropoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
167. N-(5-isopropyl-1,3-thiazol-2-yl)-1,3-benzodioxole-5-carboxamide
168. 4-(benzyloxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
169. 4-(2-cyclohexen-1-yloxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
170. N-(5-isopropyl-1,3-thiazol-2-yl)-4(trifluoromethoxy)benzamide
171. 4-(difluoromethoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
172. N-(5-isopropyl-1,3-thiazol-2-yl)-4(methylsulfanyl)benzamide
173. 2-[(4-chlorophenyl)sulfinyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
174. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[(4-nitrophenyl)sulfinyl]benzamide
175. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(4-methylphenyl)sulfonyl]-3-nitrobenzamide
176. N-(5-isopropyl-1,3-thiazol-2-yl)-3-[(trifluoromethyl)sulfanyl]benzamide
177. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxy-4-(methylsulfanyl)benzamide
178. 2-[(2-cyanophenyl)sulfanyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
179. N~1~,N-1~-diethyl-3,6-difluoro-N-2~(5-isopropyl-1,3-thiazol-2-yl)phthalamide

180. 4-formyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
181. 2-formyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
182. 4-[(2,5-dimethoxyanilino)carbonyl]amino}-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
183. 4-(hydroxymethyl)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
184. 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-2-nitrobenzyl acetate  
185. 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-2-nitrobenzyl 4-(acetylamino)-3-iodobenzoate  
186. 4-(acetylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
187. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(2-phenylacetyl)amino]benzamide  
188. 4-(acetylamino)-3-iodo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
189. 4-amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
190. 4-(dimethylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
191. 3-(dimethylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
192. 2-(methylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
193. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(trifluoromethyl)anilino]benzamide  
194. 3-[(5-bromo-1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)methyl]amino}-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
195..N-(5-isopropyl-1,3-thiazol-2-yl)-4-(1H-pyrrol-1-yl)benzamide  
196. 2,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)isonicotinamide  
197. 2-(4-bromophenyl)-6-(4-iodophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)isonicotinamide.  
198. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(trifluoromethyl)anilino]nicotinamide  
199. 2,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide  
200. 5,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide  
201. 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-6-methylnicotinamide

202. 2,6-dichloro-5-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide

203. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenoxy nicotinamide

204. N-(5-isopropyl-1,3-thiazol-2-yl)-6-(2,2,2-trifluoroethoxy)nicotinamide

205. N-(5-isopropyl-1,3-thiazol-2-yl)-2,6-dimethoxy nicotinamide

206. N-(5-isopropyl-1,3-thiazol-2-yl)-2-quinoxalinecarboxamide

207. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-pyrazinecarboxamide

208. N-(5-isopropyl-1,3-thiazol-2-yl)-8-quinolinecarboxamide

209. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-4-quinolinecarboxamide

210. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-1H-pyrazole-4-carboxamide

211. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-1H-pyrazole-3-carboxamide

212. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-pyrazole-4-carboxamide

213. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-phenyl-2H-1,2,3-triazole-4-carboxamide

214. 2-[(2,1,3-benzoxadiazol-5-yloxy)methyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-1,3-thiazole-5-carboxamide

215. N-(5-isopropyl-1,3-thiazol-2-yl)-9H-fluorene-1-carboxamide

216. N-(5-isopropyl-1,3-thiazol-2-yl)-7-methoxy-1-benzofuran-2-carboxamide

217. N-(5-isopropyl-1,3-thiazol-2-yl)-1-[(4-methylphenyl)sulfonyl]-1H-pyrrole-3-carboxamide

218. 2-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamide

219. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamide

220. N-(5-isopropyl-1,3-thiazol-2-yl)-2-naphthamide

221. N-(5-isopropyl-1,3-thiazol-2-yl)-9,10-dioxo-9,10-dihydro-2-anthracene carboxamide

222. N-(5-isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluorene-4-carboxamide

223. N-(5-isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluorene-1-carboxamide

224. N-(5-isopropyl-1,3-thiazol-2-yl)-8-oxo-5,6,7,8-tetrahydro-2-naphthalenecarboxamide

225. N-(5-isopropyl-1,3-thiazol-2-yl)-1,3-dioxo-1,3-dihydro-2-benzofuran-5-carboxamide

226. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-5-carboxamide

227. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-4-carboxamide

228. N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-2-phenyl-1H-indole-5-carboxamide

229. 2-butyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indole-5-carboxamide

230. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-6-carboxamide

231. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methoxy-1H-indole-2-carboxamide

232. 1-allyl-2-butyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-5-carboxamide

233. N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indole-2-carboxamide

234. 1-benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-1H-indole-5-carboxamide

235. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-1,2,3-benzotriazole-5-carboxamide

236. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-isoxazolecarboxamide

237. N-(5-isopropyl-1,3-thiazol-2-yl)-3-thiophenecarboxamide

238. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-thiophenecarboxamide

239. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-thiophenecarboxamide

240. 5-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-2-thiophenecarboxamide

241. N-(5-isopropyl-1,3-thiazol-2-yl)-3-[(2,3,3-trichloroacryloyl)amino]-2-thiophenecarboxamide

242. 5-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide

243. N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide

244. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(4-nitrophenyl)-2-furamide

245. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(2-nitrophenyl)-2-furamide

246. 5-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide

247. N-(5-isopropyl-1,3-thiazol-2-yl)-5-[3-(trifluoromethyl)phenyl]-2-furamide

248. 5-(4-chloro-2-nitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide

249. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(4-methyl-2-nitrophenyl)-2-furamide

250. 5-[2-chloro-5-(trifluoromethyl)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide

251. tert-butyl (1*R*)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethylcarbamate

252. (1*R*)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate

253. (1*S*)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate

254. (R,S)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

255. (R)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

256. (S)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

257. 2-(acetylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

258. (R,S)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

259. (R)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

260. (S)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

261. 3,3,3-trifluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxy-2-phenylpropanamide

262. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(1-naphthyl)acetamide

263. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-naphthyl)acetamide

264. 2-(1*H*-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

265. 2-(1,3-benzodioxol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

266. 2-(2,4-dinitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

267. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-methyl-1*H*-indol-3-yl)acetamide

268. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(1-methyl-1*H*-indol-3-yl)acetamide

269. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(5-methoxy-1*H*-indol-3-yl)acetamide

270. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(5-benzyloxy-1*H*-indol-3-yl)acetamide

271. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxy-2-methyl-1H-indol-3-yl)acetamide

272. 2-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxoacetamide

273. 2-(5-bromo-1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

274. 2-(5-fluoro-1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

275. 2-[1-(4-chlorobenzoyl)-5-methoxy-2-methyl-1H-indol-3-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

276. 3-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

277. 4-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)butanamide

278. N-(5-isopropyl-1,3-thiazol-2-yl)-3-(2-thienyl)propanamide

279. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-thienyl)acetamide

280. N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxo-2-(2-thienyl)acetamide

281. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-thienyl)acetamide

282. 2-(5-chloro-1-benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

283. 2-(1-benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

284. 2-[2-(formylamino)-1,3-thiazol-4-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamide

285. 2-{2-[(2-chloroacetyl)amino]-1,3-thiazol-4-yl}-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamide

286. 2-chloro-N-(4-(2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl)-1,3-thiazol-2-yl)acetamide

287. ethyl 2-({[2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-(1H-pyrazol-3-yl)ethylidene]amino}oxy)acetate

288. 2-(2-furyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxoacetamide

289. 2-(5-bromo-3-pyridinyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

290. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(7-methoxy-2-oxo-2H-chromen-4-yl)acetamide  
291. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenyl-3-butenamide  
292. N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxo-4-(4-methyl-phenyl)butanamide  
293. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-nitrophenyl)butanamide  
294. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenylbutanamide  
295. benzyl 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]-4-oxobutylcarbamate  
296. methyl 5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate  
297. 4-(1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)butanamide  
298. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-methoxy-1-naphthyl)-4-oxobutanamide  
299. 3-(2-chlorophenoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide  
300. 3-(4-methylphenoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide  
301. 3-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide  
302. 3-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide  
303. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylpentanamide  
304. 3-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide  
305. 3-(4-methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide  
306. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide  
307. 3-phenyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide  
308. 2-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
309. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methylbutanamide  
310. N-(5-isopropyl-1,3-thiazol-2-yl)-5-oxo-5-phenylpentanamide  
311. 2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoyl-phenylethyl acetate  
312. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[4-(1-oxo-1,3-dihydro-2H-isoindol-2-yl)phenyl]propanamide

313. 1-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)cyclopentanecarboxamide  
314. 1-phenyl-N-(5-isopropyl-1,3-thiazol-2-yl)cyclopentanecarboxamide  
315. 2-(3-bromo-4-methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
316. 2-(2-nitro-4-trifluoromethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
317. 5-cyclohexyl 1-(4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}benzyl) (2S)-2-[(tert-butoxycarbonyl)amino]pentanedioate  
318. 2-(5,6-dimethyl-1H-benzimidazol-1-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
319. 2-[5-(4-chlorophenyl)-2H-1,2,3,4-tetraazol-2-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
320. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[5-(1-pyrrolidinyl)-2H-1,2,3,4-tetraazol-2-yl]acetamide  
321. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methyl-1-benzothiophen-2-yl)acetamide  
322. N-(5-isopropyl-1,3-thiazol-2-yl)-4,4-bis(4-methylphenyl)-3-butenamide  
323. 2-cyclopropyl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
324. N-{4-bromo-6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexyl}benzamide  
325. 2-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
326. benzyl 6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexylcarbamate  
327. N-1-(5-isopropyl-1,3-thiazol-2-yl)-N-4-(2-propynyl)-2-butenediamide  
328. 4-(2,4-dimethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide  
329. 4-(4-benzyloxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide  
330. 4-(thiphen-2-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide  
331. benzyl 2-{{(benzyloxy)carbonyl}amino}-5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate  
332. 4-(1H-indol-3-yl)-N-{3-[(5-isopropyl-1,3-thiazol-2-yl)amino]-3-oxopropyl}butanamide

333. 4-{{(5-isopropyl-1,3-thiazol-2-yl)amino}carbonyl}phenyl 4-chlorobenzencesulfonate

334. N-(5-isopropyl-1,3-thiazol-2-yl)-4-{{(2-methoxyanilino)carbonyl}amino}benzamide

335. 4-{{[2-(isopropylsulfonyl)acetyl]amino}-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

336. N-(5-isopropyl-1,3-thiazol-2-yl)-4-{{[2-(phenylsulfonyl)acetyl]amino}benzamide

337. 4-[(diethylamino)sulfonyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

338. 2-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

339. 3,5-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

340. 3-{{(2-fluoroanilino)carbonyl}amino}-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

341. N-(5-isopropyl-1,3-thiazol-2-yl)-1-phenyl-5-propyl-1H-pyrazole-4-carboxamide

342. 3-chloro-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfonyl)-2-thiophenecarboxamide

343. 3-iodo-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfonyl)-2-thiophenecarboxamide

344. 2-{{(4-chlorophenyl)sulfonyl)methyl}-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-1,3-thiazole-5-carboxamide

345. 5-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(trifluoromethyl)-3-furamide

346. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,3,4,5,6-pentafluorophenyl)acetamide

347. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-fluorophenyl)acetamide

348. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-bromophenyl)acetamide

349. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-chlorophenyl)acetamide

350. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-nitrophenyl) acetamide

351. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-trifluoromethylphenyl)acetamide

352. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-methoxyphenyl)acetamide

353. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-dimethoxyphenyl)acetamide

354. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-difluorophenyl)acetamide  
355. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4,5-trimethoxyphenyl)acetamide  
356. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,6-dichlorophenyl)acetamide  
357. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-chloro-6-fluorophenyl)acetamide  
358. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,5-dimethoxyphenyl)acetamide  
359. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,5-difluorophenyl)acetamide  
360. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-bistrifluoromethylphenyl)acetamide  
361. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methylthiophenyl)acetamide  
362. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide  
363. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-bromophenyl)acetamide  
364. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-chlorophenyl)acetamide  
365. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-fluorophenyl)acetamide  
366. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl)acetamide  
367. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-trifluoromethylphenyl)acetamide  
368. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methylphenyl)acetamide  
369. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-dimethylaminophenyl)acetamide  
370. 2-[1,1'-biphenyl]-4-yl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide,  
371. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-trifluoromethylphenyl)acetamide  
372. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-bromophenyl)acetamide  
373. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-chlorophenyl)acetamide  
374. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-nitrophenyl)acetamide  
375. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide  
376. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-dinitrophenyl)acetamide  
377. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-dichlorophenyl)acetamide

378. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-difluorophenyl)acetamide  
379. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-benzyloxy-3-methoxyphenyl)acetamide  
380. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dichlorophenyl)acetamide  
381. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-difluorophenyl)acetamide  
382. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dimethoxyphenyl)acetamide  
383. 2-(2,3-dihydro-1H-inden-5-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
384. N-(5-isopropyl-1,3-thiazol-2-yl)-1-phenylcyclopropanecarboxamide  
385. 2-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide  
386. 2-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide  
387. N-(5-isopropyl-1,3-thiazol-2-yl)-2,2-diphenylacetamide  
388. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-nitrophenoxy)acetamide  
389. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl)propanamide  
390. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide  
391. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-isobutylphenyl)propanamide  
392. N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxo-2-phenylacetamide  
393. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-phenylpentanamide  
394. (E, Z)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-2-butenamide  
395. N-(5-isopropyl-1,3-thiazol-2-yl)bicyclo[4.2.0]octa-1,3,5-triene-7-carboxamide  
396. N-(5-isopropyl-1,3-thiazol-2-yl)-3-oxo-1-indanecarboxamide  
397. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylbutanamide  
398. tert-butyl (1S)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-1-methyl-2-oxoethylcarbamate  
399. tert-butyl (1S,2S)-1-{{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-2-methylbutylcarbamate  
400. tert-butyl 2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

401. *tert*-butyl (1*S*)-5-amino-1-{{(5-isopropyl-1,3-thiazol-2-yl)amino}carbonyl}pentylcarbamate

402. *tert*-butyl 4-[(imino{[(4-methylphenyl)sulfonyl]amino}methyl)amino]-1-{{(5-isopropyl-1,3-thiazol-2-yl)amino}carbonyl}butylcarbamate

403. *tert*-butyl 1-{{(5-isopropyl-1,3-thiazol-2-yl)amino}carbonyl}-3-(tritylamino)propylcarbamate

404. *tert*-butyl (1*S*)-1-(benzyloxyethyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

405. *tert*-butyl (1*S*)-1-benzyl-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

406. *tert*-butyl (1*R*)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-(benzylthiomethyl)ethylcarbamate.

407. benzyl (3*S*)-3-[(*tert*-butoxycarbonyl)amino]-4-[(5-isopropyl-1,3-thiazol-2-yl)amino]-4-oxobutanoate

408. *tert*-butyl (2*S*)-2-{{(5-isopropyl-1,3-thiazol-2-yl)aminolcarbonyl}-1-pyrrolidinocarboxylate

409. *tert*-butyl (1*S*)-1-(1*H*-indol-3-ylmethyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

410. *tert*-butyl (1*S*)-1-{{(5-isopropyl-1,3-thiazol-2-yl)amino}carbonyl}-3-(methylsulfanyl)propylcarbamate

411. *tert*-butyl (1*S*)-2-benzyloxy-1-{{(5-isopropyl-1,3-thiazol-2-yl)amino}carbonyl}propylcarbamate

412. *tert*-butyl (1*S*)-1-(4-benzyloxybenzyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

413. tert-butyl (1*S*)-1-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-2-methylpropylcarbamate

414. tert-butyl (1*S*)-1-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-3-methylbutylcarbamate

415. benzyl (4*S*)-4-[(tert-butoxycarbonyl)amino]-5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate

- 1 -

2-AMINO-THIAZOLE DERIVATIVES, PROCESS FOR THEIR PREPARATION, AND THEIR USE AS ANTITUMOR AGENTS.

5

The present invention relates to 2-amino-thiazole derivatives, to a process for their preparation, to pharmaceutical compositions containing them and to their use as therapeutic agents, particularly in the treatment of 10 cancer and cell proliferative disorders.

Several cytotoxic drugs such as, e.g. fluorouracil (5-FU), doxorubicin and camptothecins result to damage DNA or to affect cellular metabolic pathways and thus cause, in many 15 cases, an indirect block of the cell cycle.

Therefore, by producing an irreversible damage to both normal and tumor cells, these agents result in a significant toxicity and side-effects.

In this respect, compounds capable of being highly specific 20 antitumor agents by selectively leading to tumor cell arrest and apoptosis, with comparable efficacy but reduced toxicity than the currently available drugs, are desirable.

It is well known in the art that progression through the 25 cell cycle is governed by a series of checkpoint controls, otherwise referred to as restriction points, which are regulated by a family of enzymes known as the cyclin-dependent kinases (cdk).

In their turn, the cdk's themselves are regulated at many 30 levels such as, for instance, binding to cyclins.

A normal progression through the cell cycle is controlled by the coordinated activation and inactivation of different cyclin/cdk complexes. In G1, both cyclin D/cdk4 and cyclin E/cdk2 are thought to mediate the onset of S-phase. 35 Progression through S-phase requires the activity of cyclin A/cdk2 whereas the activation of cyclin A/cdc2 (cdk1) and cyclin B/cdc2 are required for the onset of metaphases.

For a general reference to cyclins and cyclin-dependent kinases see, for instance, Kevin R. Webster et al. in *Exp. Opin. Invest. Drugs*, 1998, Vol. 7(6), 865-887.

5 Checkpoint controls are defective in tumor cells due, in part, to disregulation of cdk activity. For example, altered expression of cyclin E and cdk's has been observed in tumor cells, and deletion of the cdk inhibitor p27 KIP gene in mice has been shown to result in a higher incidence  
10 of cancer.

Increasing evidence supports the idea that the cdk's are rate-limiting enzymes in cell cycle progression and, as such, represent molecular targets for therapeutic intervention. In particular, the direct inhibition of  
15 cdk/cyclin kinase activity should be helpful in restricting the unregulated proliferation of a tumor cell.

It has now been found that the 2-amino-1,3-thiazoles of the invention are endowed with cdk/cyclin kinase inhibitory activity and are thus useful in therapy as antitumor agents whilst lacking, in terms of both toxicity and side effects, the aforementioned drawbacks known for currently available  
20 antitumor drugs.

More specifically, the compounds of this invention are  
25 useful in the treatment of a variety of cancers including, but not limited to: carcinoma such as bladder, breast, colon, kidney, liver, lung, including small cell lung cancer, esophagus, gall-bladder, ovary, pancreas, stomach, cervix, thyroid, prostate, and skin, including squamous  
30 cell carcinoma; hematopoietic tumors of lymphoid lineage, including leukemia, acute lymphocytic leukemia, acute lymphoblastic leukemia, B-cell lymphoma, T-cell-lymphoma, Hodgkin's lymphoma, non-Hodgkin's lymphoma, hairy cell lymphoma and Burkett's lymphoma; hematopoietic tumors of  
35 myeloid lineage, including acute and chronic myelogenous leukemias, myelodysplastic syndrome and promyelocytic leukemia; tumors of mesenchymal origin, including fibrosarcoma and rhabdomyosarcoma; tumors of the central

and peripheral nervous system, including astrocytoma, neuroblastoma, glioma and schwannomas; other tumors, including melanoma, seminoma, teratocarcinoma, osteosarcoma, xenoderoma pigmentosum, keratoctanthoma, 5 thyroid follicular cancer and Kaposi's sarcoma.

Due to the key role of cdk5 in the regulation of cellular proliferation, these 2-amino-1,3-thiazole derivatives are also useful in the treatment of a variety of cell 10 proliferative disorders such as, for instance, benign prostate hyperplasia, familial adenomatosis, polyposis, neuro-fibromatosis, psoriasis, vascular smooth cell proliferation associated with atherosclerosis, pulmonary fibrosis, arthritis, glomerulonephritis and post-surgical 15 stenosis and restenosis.

The compounds of the invention can be useful in the treatment of Alzheimer's disease, as suggested by the fact that cdk5 is involved in the phosphorylation of tau protein (J. Biochem., 117, 741-749, 1995).

20 The compounds of this invention, as modulators of apoptosis, could be useful in the treatment of cancer, viral infections, prevention of AIDS development in HIV-infected individuals, autoimmune diseases and neurodegenerative disorder.

25 The compounds of this invention could be useful in inhibiting tumor angiogenesis and metastasis.

The compounds of this invention may also act as inhibitors 30 of other protein kinases, e.g. protein kinase C, her2, raf1, MEK1, MAP kinase, EGF receptor, PDGF receptor, IGF receptor, PI3 kinase, weel kinase, Src, Abl and thus be effective in the treatment of diseases associated with other protein kinases.

35 Several 2-amino-1,3-thiazole derivatives are known in the art. Just few examples among them are 2-acetamido-, 2-propionamido- or 2-butyramido-1,3-thiazole derivatives further substituted by halogen atoms in position 5 of the

thiazole ring, which are reported as herbicides in JP 73027467 (Sankyo Co. Ltd.) or US 3,374,082 (The Upjohn Co.); 5-nitro-2-benzamido-1,3-thiazole is reported as pesticide in Ann. Rech. Vet., 22(4), 359-63, 1991; 5-phenyl-2-acetamido-1,3-thiazoles further substituted onto phenyl ring are reported as synthetic intermediates (Chemical Abstracts, 1980, 92:128793); and 5-dimethylaminomethyl- or 5-diethylaminomethyl-2-acetamido-1,3-thiazole, both reported as herbicides in JP 71018564 (Japan Gas Chem Co.).

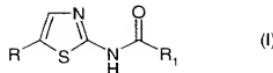
Several other 2-amino-1,3-thiazole derivatives have been reported in the art as useful therapeutic agents.

In particular, 5-methyl-1,3-thiazoles further substituted in position 2 of the thiazole ring by a benzothiazinyl-carbonylamino moiety, or derivatives thereof, have been described as cyclooxygenase inhibitors; see, for instance, C.A. 126(1997):301540.

2-Benzamido-1,3-thiazoles are disclosed in EP-A-261503 (Valeas S.p.A.) as antiallergic agents; 5-Alkyl-2-phenylalkylcarbonylamino-1,3-thiazoles further substituted onto the phenyl ring with an alkenylcarbonyl or alkynylcarbonyl moieties are disclosed in WO 98/04536 (Otsuka Pharmaceutical Co.) as protein kinase C inhibitors. 5-Arylthio-2-acylamino-1,3-thiazole derivatives are disclosed in EP-A-412404 (Fujisawa Pharm. Co.) as antitumor agents.

In addition, among the compounds reported in the art as therapeutic agents, DE 2128941 (Melle-Bezons) discloses 2-aminomethylcarbonylamino-5-chloro-1,3-thiazoles as antiinflammatory, sedative and analgesic agents; the compound 2-diethylaminomethylcarbonylamino-5-chloro-1,3-thiazole being specifically exemplified therein.

Accordingly, the present invention provides the use of a compound which is a 2-amino-1,3-thiazole derivative of formula (I)



wherein

R is a halogen atom, a nitro group, an optionally substituted amino group or it is a group, optionally further substituted, selected from:

- i) straight or branched C<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>2</sub>-C<sub>6</sub> alkenyl or C<sub>2</sub>-C<sub>6</sub> alkynyl;
- ii) C<sub>3</sub>-C<sub>6</sub> cycloalkyl;
- iii) aryl or arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

R<sub>1</sub> is an optionally further substituted group selected from:

- i) straight or branched C<sub>1</sub>-C<sub>6</sub> alkyl or C<sub>2</sub>-C<sub>6</sub> alkenyl;
- ii) 3 to 6 membered carbocycle or 5 to 7 membered heterocycle ring;
- iii) aryl or arylcarbonyl;
- iv) arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;
- v) arylalkenyl with from 2 to 6 carbon atoms within the straight or branched alkenyl chain;
- vi) an optionally protected amino acid residue; or a pharmaceutically acceptable salt thereof; in the manufacture of a medicament for treating cell proliferative disorders associated with an altered cell dependent kinase activity.

According to a preferred embodiment of the invention, the said cell proliferative disorder is selected from the group consisting of cancer, Alzheimer's disease, viral infections, auto-immune diseases or neurodegenerative disorders.

Preferably, the cancer is selected from the group consisting of carcinoma, squamous cell carcinoma, hematopoietic tumors of myeloid or lymphoid lineage, tumors of mesenchymal origin, tumors of the central and peripheral nervous system, melanoma, seminoma, teratocarcinoma,

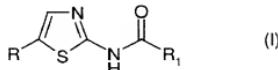
osteosarcoma, xenoderoma pigmentosum, keratoctanthoma, thyroid follicular cancer and Kaposi's sarcoma.

According to another preferred embodiment of the invention, 5 the cell proliferative disorder is selected from the group consisting of benign prostate hyperplasia, familial adenomatosis polyposis, neuro-fibromatosis, psoriasis, vascular smooth cell proliferation associated with atherosclerosis, pulmonary fibrosis, arthritis 10 glomerulonephritis and post-surgical stenosis and restenosis.

In addition, being useful in the treatment of cell proliferative disorders associated with an altered cell 15 dependent kinase activity, hence cell cycle inhibition or cdk/cyclin dependent inhibition, the compounds of formula (I) of the invention also enable tumor angiogenesis and metastasis inhibition.

20 As above reported, some of the compounds of formula (I) of the invention have been reported in the art as useful therapeutic agents, for instance as antiinflammatory, sedative and analgesic agents.

25 Therefore, it is a further object of the present invention a compound which is a 2-amino-1,3-thiazole derivative of formula (I)



wherein

30 R is a halogen atom, a nitro group, an optionally substituted amino group or it is a group, optionally further substituted, selected from:

- i) straight or branched C<sub>1</sub>-C<sub>8</sub> alkyl, C<sub>2</sub>-C<sub>6</sub> alkenyl or C<sub>2</sub>-C<sub>6</sub> alkynyl;
- 35 ii) C<sub>3</sub>-C<sub>6</sub> cycloalkyl;

iii) aryl or arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

R<sub>1</sub> is an optionally further substituted group selected from:

5 i) straight or branched C<sub>1</sub>-C<sub>6</sub> alkyl or C<sub>2</sub>-C<sub>6</sub> alkenyl;

ii) 3 to 6 membered carbocycle or 5 to 7 membered heterocycle ring;

iii) aryl or arylcarbonyl;

iv) arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

10 v) arylalkenyl with from 2 to 6 carbon atoms within the straight or branched alkenyl chain;

vi) an optionally protected amino acid residue;

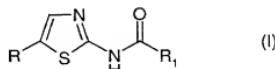
or a pharmaceutically acceptable salt thereof; for use as a

15 medicament; provided that each of R and R<sub>1</sub>, independently, is not a methyl group and that the compound is not 2-diethylaminomethyl-carbonylamino-5-chloro-1,3-thiazole.

Among the compounds of formula (I) above reported, several

20 derivatives result to be novel.

Therefore, the present invention further provides a compound which is a 2-amino-1,3-thiazole derivative of formula (I)



25

wherein

R is a halogen atom, a nitro group, an optionally substituted amino group or it is a group, optionally further substituted, selected from:

30 i) straight or branched C<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>2</sub>-C<sub>6</sub> alkenyl or C<sub>2</sub>-C<sub>6</sub> alkynyl;

ii) C<sub>3</sub>-C<sub>6</sub> cycloalkyl;

iii) aryl or arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

35 R<sub>1</sub> is an optionally further substituted group selected from:

i) straight or branched C<sub>1</sub>-C<sub>8</sub> alkyl or C<sub>2</sub>-C<sub>6</sub> alkenyl;

ii) 3 to 6 membered carbocycle or 5 to 7 membered heterocycle ring;

iii) aryl or arylcarbonyl;

5 iv) arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

v) arylalkenyl with from 2 to 6 carbon atoms within the straight or branched alkenyl chain;

vi) an optionally protected amino acid residue;

10 or a pharmaceutically acceptable salt thereof;

provided that:

a) R and R<sub>1</sub>, each independently, are not methyl;

b) when R is bromine or chlorine then, R<sub>1</sub> is not unsubstituted C<sub>2</sub>-C<sub>4</sub> alkyl or an optionally substituted

15 aminomethyl;

c) when R is nitro or phenyl, then R<sub>1</sub> is not unsubstituted phenyl.

20 The compounds of formula (I) may have asymmetric carbon atoms and may therefore exist either as racemic admixtures or as individual optical isomers.

25 Accordingly, all the possible isomers and their admixtures and of both the metabolites and the pharmaceutically acceptable bio-precursors (otherwise referred to as prodrugs) of the compounds of formula (I), as well as the uses thereof, are also within the scope of the present invention.

30 In the present description, unless otherwise specified, with the term halogen atom we intend a chlorine, bromine, fluorine or iodine atom.

35 With the term optionally substituted amino group we intend an amino group wherein one or both hydrogen atoms are optionally replaced by other substituents which are the same or different, as set forth below.

With the term straight or branched C<sub>1</sub>-C<sub>8</sub> alkyl we intend a group such as, for instance, methyl, ethyl, n.propyl, isopropyl, n.butyl, isobutyl, sec-butyl, tert-butyl, n.pentyl, n.hexyl, n.heptyl, n.octyl and the like.

5

With the term straight or branched C<sub>2</sub>-C<sub>6</sub> alkenyl or alkynyl we intend a group such as, for instance, vinyl, allyl, isopropenyl, 1-, 2- or 3-butenyl, isobutylene, pentenyl, hexenyl, ethynyl, 1- or 2-propynyl, butynyl, pentynyl, 10 hexynyl and the like.

With the term C<sub>3</sub>-C<sub>6</sub> cycloalkyl we intend a cyclopropyl, cyclobutyl, cyclopentyl or cyclohexyl group.

15 With the term aryl, either as such or as arylalkyl, arylalkenyl, arylcarbonyl and the like, we intend a mono-, bi- or poly- either carbocyclic as well as heterocyclic hydrocarbon with from 1 to 4 ring moieties, either fused or linked to each other by single bonds, wherein at least one 20 of the carbocyclic or heterocyclic rings is aromatic.

Examples of aryl groups are phenyl, indanyl, biphenyl,  $\alpha$ - or  $\beta$ -naphthyl, fluorenyl, 9,10-dihydroanthracenyl, pyridyl, pyrazinyl, pyrimidinyl, pyridazinyl, indolyl, imidazolyl, 1,2-methylenedioxyphenyl, thiazolyl, isothiazolyl,

25 pyrrolyl, pyrrolyl-phenyl, furyl, phenyl-furyl, benzotetrahydrofuran, oxazolyl, isoxazolyl, pyrazolyl, chromenyl, thieryl, benzothienyl, isoindolinyl, benzoimidazolyl, tetrazolyl, tetrazolylphenyl, pyrrolidinyl-tetrazolyl, isoindolinyl-phenyl, quinolinyl, 30 isoquinolinyl, 2,6-diphenyl-pyridyl, quinoxalinyl, pyrazinyl, phenyl-quinolyl, benzofurazanyl, 1,2,3-triazole, 1-phenyl-1,2,3-triazole, and the like.

With the term 3 to 6 membered carbocycle, hence 35 encompassing but not limited to C<sub>3</sub>-C<sub>6</sub> cycloalkyl groups, we also intend an unsaturated carbocyclic hydrocarbon such as, for instance, cyclopentylene or cyclohexylene.

With the term 5 to 7 membered heterocycle, hence encompassing aromatic heterocycles also referred to as aryl groups, we further intend a saturated or partially unsaturated 5 to 7 membered carbocycle wherein one or more carbon atoms are replaced by heteroatoms such as nitrogen, oxygen and sulphur.

Examples of 5 to 7 membered heterocycles, optionally benzocondensed or further substituted, are 1,3-dioxolane, 10 pyran, pyrrolidine, pyrroline, imidazolidine, pyrazolidine, pyrazoline, piperidine, piperazine, N-alkyl-piperazine, morpholine, tetrahydrofuran and the like.

With the term amino acid residue we intend the residue of a natural  $\alpha$ -amino acid of formula HOOC-R<sub>1</sub>, wherein R<sub>1</sub> is bonded to the thiazole-NH-C(=O)- moiety and is represented by a -CH(Z)NHY group wherein Z is the characterising portion of the amino acid and Y is hydrogen or a suitable amino protecting group such as, for instance, tert-butoxycarbonyl or benzyloxycarbonyl.

20 Examples of  $\alpha$ -amino acids are alanine, isoleucine, glycine, lysine, arginine, cystine, histidine, leucine, proline and the like.

According to the above indicated substituent meanings, any 25 of the above R and R<sub>1</sub> groups may be optionally substituted in any of the free positions by one or more groups, for instance 1 to 6 groups, selected from: halogen, nitro, oxo groups (=O), carboxy, cyano, alkyl, perfluorinated alkyl, 30 alkenyl, alkynyl, cycloalkyl, aryl, heterocycyl; amino groups and derivatives thereof such as, for instance, alkylamino, alkoxy carbonyl alkylamino, dialkylamino, arylamino, diarylamino or arylureido; carbonylamino groups and derivatives thereof such as, for instance, hydrogenocarbonylamino (HCONH-), alkylcarbonylamino, 35 alkenylcarbonylamino, arylcarbonylamino, alkoxy carbonylamino; oxygen-substituted oximes such as, for instance, alkoxy carbonyl alkoxyimino or alkoxyimino; hydroxy

groups and derivatives thereof such as, for instance, alkoxy, aryloxy, alkylcarbonyloxy, arylcarbonyloxy, cycloalkenyloxy; carbonyl groups and derivatives thereof such as, for instance, alkylcarbonyl, arylcarbonyl, 5 alkoxy carbonyl, aryloxycarbonyl, cycloalkyloxycarbonyl, aminocarbonyl, alkylaminocarbonyl, dialkylaminocarbonyl; sulfurated derivatives such as, for instance, alkylthio, arylthio, alkylsulphonyl, arylsulphonyl, alkylsulphanyl, arylsulphanyl, arylsulphonyloxy, aminosulfonyl, 10 alkylaminosulphonyl or dialkylaminosulphonyl. In their turn, whenever appropriate, each of the above possible substituents on R and R<sub>1</sub> groups may be further substituted by one or more of the aforementioned groups.

Examples of compounds of formula (I) wherein R and R<sub>1</sub> groups 15 are substituted by one or more of the aforementioned substituents which, in their turn, are optionally further substituted as set forth above, are given below.

Pharmaceutically acceptable salts of the compounds of 20 formula (I) are the acid addition salts with inorganic or organic, e.g. nitric, hydrochloric, hydrobromic, sulphuric, perchloric, phosphoric, acetic, trifluoroacetic, propionic, glycolic, lactic, oxalic, malonic, malic, maleic, tartaric, citric, benzoic, cinnamic, mandelic, methanesulphonic, 25 isethionic and salicylic acid, as well as the salts with inorganic or organic bases, e.g. alkali or alkaline-earth metals, especially sodium, potassium, calcium or magnesium hydroxides, carbonates or bicarbonates, acyclic or cyclic amines, preferably methylamine, ethylamine, diethylamine, 30 triethylamine or piperidine.

The compounds of formula (I) may have asymmetric carbon atoms and may therefore exist either as racemic admixtures or as individual optical isomers.

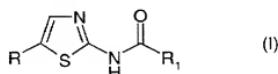
35 Accordingly, the use as an antitumor agent of all the possible isomers and their admixtures and of both the metabolites and the pharmaceutically acceptable bio-precursors (otherwise referred to as pro-drugs) of the

compounds of formula (I) are also within the scope of the present invention.

Preferred compounds of formula (I), according to the 5 present invention, are 2-amino-1,3-thiazole derivatives wherein R is a halogen atom or an optionally substituted group selected from a straight or branched C<sub>1</sub>-C<sub>4</sub> alkyl, C<sub>3</sub>-C<sub>6</sub> cycloalkyl, aryl or an arylalkyl with from 1 to 4 carbon atoms within the alkyl chain; R<sub>1</sub> is an optionally 10 substituted group selected from straight or branched C<sub>1</sub>-C<sub>4</sub> alkyl or alkenyl, aryl or arylalkyl with from 1 to 4 carbon atoms within the alkyl chain or it is an optionally protected amino acid residue.

15 Still more preferred compounds, within this class, are the derivatives of formula (I) wherein R is a bromine or chlorine atom or is an optionally substituted group selected from straight or branched C<sub>1</sub>-C<sub>4</sub> alkyl, cyclopropyl, aryl or arylalkyl with from 1 to 2 carbon atoms within the 20 alkyl chain; R<sub>1</sub> is an optionally substituted group selected from straight or branched C<sub>1</sub>-C<sub>4</sub> alkyl or alkenyl, aryl or arylalkyl with from 1 to 4 carbon atoms within the alkyl chain or it is an optionally protected amino acid residue.

25 Another class of preferred compounds of the invention are the compounds of formula (I)



wherein

30 R is a halogen atom or is selected from nitro, amino, alkylamino, hydroxylalkylamino, arylamino, C<sub>3</sub>-C<sub>6</sub> cycloalkyl and straight or branched C<sub>1</sub>-C<sub>6</sub> alkyl which is unsubstituted or substituted by hydroxy, alkylthio, alkoxy, amino, alkylamino, alkoxy carbonylamino, alkoxy carbonylalkylamino, alkylcarbonyl, alkylsulfonyl, 35 alkoxy carbonyl, carboxy or aryl which is unsubstituted

or substituted by one or more hydroxy, halogen, nitro, alkoxy, aryloxy, alkylthio, arylthio, amino, alkylamino, dialkylamino, N-alkyl-piperazinyl, 4-morpholinyl, arylamino, cyano, alkyl, phenyl, aminosulfonyl, aminocarbonyl, alkylcarbonyl, arylcarbonyl, alkoxycarbonyl or carboxy groups, or R is an aryl group which is unsubstituted or substituted by one or more hydroxy, halogen, nitro, alkoxy, aryloxy, alkylthio, arylthio, amino, alkylamino, dialkylamino, N-alkyl-piperazinyl, 4-morpholinyl, arylamino, cyano, alkyl, phenyl, aminosulfonyl, aminocarbonyl, alkylcarbonyl, arylcarbonyl, alkoxycarbonyl or carboxy groups;

R<sub>1</sub> is a straight or branched C<sub>1</sub>-C<sub>6</sub> alkyl group or an aryl group, each being unsubstituted or substituted as defined above for R;

or a pharmaceutically acceptable salt thereof;

provided that:

- a) R and R<sub>1</sub>, each independently, are not methyl;
- b) when R is bromine or chlorine then, R<sub>1</sub> is not unsubstituted C<sub>2</sub>-C<sub>4</sub> alkyl or an optionally substituted aminomethyl;
- c) when R is nitro or phenyl, then R<sub>1</sub> is not unsubstituted phenyl.

25

Examples of preferred compounds of the invention, whenever appropriate in the form of pharmaceutically acceptable salts, e.g. hydrobromide or hydrochloride salt, are the following:

- 30 1. ethyl 3-[(5-bromo-1,3-thiazol-2-yl)amino]-3-oxopropanoate;
2. N-(5-bromo-1,3-thiazol-2-yl)-2-phenyl-acetamide;
3. N-(5-bromo-1,3-thiazol-2-yl)-benzamide;
4. Ethyl 4-[(5-bromo-1,3-thiazol-2-yl)amino]-4-oxobutanoate;
- 35 5. N-(5-Bromo-thiazol-2-yl)-3-hydroxy-propionamide;
6. N-(5-Bromo-1,3-thiazol-2-yl)-4-hydroxybutanamide;

7. N-(5-Bromo-thiazol-2-yl)-2-ethoxy-acetamide;

8. 2-N-[2-(3-pyridyl)-acetyl-amino]-5-bromo-thiazole;

9. 2-N-[2-(3-pyridyl)-acetyl-amino]-5-isopropyl-thiazole;

10. N-(5-bromo-1,3-thiazol-2-yl)-2-(3-hydroxyphenyl)acetamide;

11. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-hydroxyphenyl)acetamide

12. N-(5-bromo-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide;

13. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide;

14. N-(5-bromo-1,3-thiazol-2-yl)-2-(3-chorophenyl)acetamide;

15. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-chorophenyl)acetamide;

16. N-(5-bromo-1,3-thiazol-2-yl)-2-(4-hydroxyphenyl)acetamide;

17. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxyphenyl)acetamide;

18. N-(5-bromo-1,3-thiazol-2-yl)-2-(3,4-dihydroxyphenyl)acetamide;

19. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dihydroxyphenyl)acetamide;

20. N-(5-bromo-1,3-thiazol-2-yl)-2-(4-hydroxy-3-methoxyphenyl)acetamide;

21. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxy-3-methoxyphenyl)acetamide;

22. N-(5-bromo-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide;

23. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide;

24. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-chlorophenyl)acetamide;

25. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-acetamide;

36. N-(5-bromo-thiazol-2-yl)-4-sulfamoyl-benzamide;

27. N-(5-isopropyl-thiazol-2-yl)-4-sulfamoyl-benzamide;

28. 4-amino-N-(5-bromo-1,3-thiazol-2-yl)butanamide;

29. 3-amino-N-(5-bromo-1,3-thiazol-2-yl)propionamide;  
30. N-(5-isopropyl-1,3-thiazol-2-yl)-butanamide;  
31. N-(5-bromo-1,3-thiazol-2-yl)-butanamide;  
32. N-(5-chloro-1,3-thiazol-2-yl)-butanamide;  
5 33. N-(5-phenyl-1,3-thiazol-2-yl)-butanamide;  
34. N-(5-nitro-1,3-thiazol-2-yl)-butanamide;  
35. N-(5-methyl-1,3-thiazol-2-yl)-butanamide;  
36. N-(5-benzyl-1,3-thiazol-2-yl)-butanamide;  
37. N-(5-isobutyl-1,3-thiazol-2-yl)-butanamide;  
10 38. N-(5-cyclopropyl-1,3-thiazol-2-yl)-butanamide;  
39. N-{5-[2-(methylsulfonyl)ethyl]-1,3-thiazol-2-yl}-butanamide;  
40. N-[5-(2-methylthioethyl)-1,3-thiazol-2-yl]-butanamide;  
41. N-{5-[2-(methoxycarbonyl)ethyl]-1,3-thiazol-2-yl}-butanamide;  
15 42. N-[5-(3-methoxy-propyl)-1,3-thiazol-2-yl]-butanamide;  
43. N-[5-(2-ethoxy-ethyl)-1,3-thiazol-2-yl]-butanamide;  
44. N-[5-(indol-3-yl-methyl)-1,3-thiazol-2-yl]-butanamide;  
45. N-[5-(3-oxo-butyl)-1,3-thiazol-2-yl]-butanamide;  
20 46. 2-[3-(3-chloropropoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
47. 2-[3-(2-chloroethoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
48. 2-(4-aminophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
25 49. 4-amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
50. 2-(2-amino-1,3-thiazol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
51. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-[3-(4-morpholinyl)propoxy]phenyl)acetamide  
30 52. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-[2-(4-morpholinyl)ethoxy]phenyl)acetamide  
53. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-[3-(1-pyrrolidinyl)propoxy]phenyl)acetamide  
35 54. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-[3-(4-methyl-1-piperazinyl)propoxy]phenyl)acetamide

55. 2-[3-[2-(dimethylamino)ethoxy]phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;

56. 2-[3-[3-(dimethylamino)propoxy]phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

5 57. 2-[4-(dimethylamino)phenyl]-N-(5-isobutyl-1,3-thiazol-2-yl)acetamide

58. 2-(1,3-benzodioxol-5-yl)-N-(5-isobutyl-1,3-thiazol-2-yl)acetamide

59. N-(5-benzyl-1,3-thiazol-2-yl)-2-[4-(dimethylamino)phenyl]acetamide

10 60. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(2-methoxyethoxy)-phenyl]acetamide

61. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-methyl-1-piperazinyl)benzamide

15 62. N-(5-isobutyl-1,3-thiazol-2-yl)-2-(3-pyridinyl)acetamide

63. N-(5-benzyl-1,3-thiazol-2-yl)-2-(3-pyridinyl)acetamide

64. 2-[N-[2'-N'-(ethoxycarbonyl-methyl)-amino]-acetyl]-amino-5-bromo-thiazole

20 65. 2-anilino-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

66. (R)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide

67. (S)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide

25 68. N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

69. 2,5-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

70. 3,5-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

71. 3,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

30 72. 2,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

73. 2,3-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

74. 3-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

75. 2-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

76. 4-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

35 77. 3-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

78. 4-chloro-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

79. 5-bromo-2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
80. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
81. 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
5 82. 4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
83. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
84. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
85. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
86. 2,4-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
10 87. 3,4-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
88. 2,3,4,5,6-pentafluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
89. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-3-nitrobenzamide  
15 90. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-nitrobenzamide  
91. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-nitrobenzamide  
92. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-nitrobenzamide  
20 93. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxy-2-nitrobenzamide  
94. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-2-nitrobenzamide  
25 95. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxy-3-nitrobenzamide  
96. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-nitrobenzamide  
97. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dinitrobenzamide  
30 98. 5-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl-2-nitrophenyl octanoate  
99. N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide  
100. N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide  
101. N-(5-isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamide  
35 102. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl)-3-nitrobenzamide  
103. 4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide

104. 6-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide

105. 4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide

5 106. 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamide

107. 5-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide

108. 2-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-5-nitrobenzamide

10 109. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide

110. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide

15 111. N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitro-4-(trifluoromethyl)benzamide

112. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-bis(trifluoromethyl)benzamide

113. N-(5-isopropyl-1,3-thiazol-2-yl)-2,6-bis(trifluoromethyl)benzamide

20 114. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(trifluoromethyl)benzamide

115. N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide

25 116. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethyl)benzamide

117. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide

118. 5-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide

30 119. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethyl)benzamide

120. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide

35 121. methyl 4-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}benzoate

122. methyl 2-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}benzoate

123. 4-cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
124. 3-cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
125. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methylbenzamide  
126. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methylbenzamide  
5 127. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylbenzamide  
128. N-(5-isopropyl-1,3-thiazol-2-yl)-4-vinylbenzamide  
129. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(2-  
phenylethynyl)benzamide  
130. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-  
10 methylbenzamide  
131. 2-benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
132. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenethylbenzamide  
133. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylbenzamide  
134. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenylbenzamide  
15 135. 4-(tert-butyl)-N-(5-isopropyl-1,3-thiazol-2-  
yl)benzamide  
136. N-(5-isopropyl-1,3-thiazol-2-yl)-4-isopropylbenzamide  
137. N-(5-isopropyl-1,3-thiazol-2-yl)-4-pentylbenzamide  
138. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-  
20 methylbenzamide  
139. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4-dimethylbenzamide  
140. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethylbenzamide  
141. 4-acetyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
142. N-(5-isopropyl-1,3-thiazol-2-yl)-4-  
25 (methylsulfonyl)benzamide  
143. 5-(aminosulfonyl)-2,4-dichloro-N-(5-isopropyl-1,3-  
thiazol-2-yl)benzamide  
144. 5-(aminosulfonyl)-4-chloro-N-(5-isopropyl-1,3-thiazol-  
2-yl)benzamide  
30 145. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-  
methoxybenzamide  
146. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-  
methoxybenzamide  
147. 5-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-  
35 methoxybenzamide  
148. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide  
149. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxybenzamide  
150. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxybenzamide

151. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4-dimethoxybenzamide

152. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethoxybenzamide

5 153. N-(5-isopropyl-1,3-thiazol-2-yl)-2,4-dimethoxybenzamide

154. N-(5-isopropyl-1,3-thiazol-2-yl)-2,3-dimethoxybenzamide

155. N-(5-isopropyl-1,3-thiazol-2-yl)-3-phenoxybenzamide

10 156. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenoxybenzamide

157. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenoxybenzamide

158. 2-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

159. 4-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

160. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4,5-trimethoxybenzamide

15 161. 3,4-diethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

162. 3,4,5-triethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

163. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-(methoxymethoxy)benzamide

20 164. 4-butoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

165. N-(5-isopropyl-1,3-thiazol-2-yl)-4-propoxybenzamide

166. 4-isopropoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

167. N-(5-isopropyl-1,3-thiazol-2-yl)-1,3-benzodioxole-5-carboxamide

25 168. 4-(benzyloxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

169. 4-(2-cyclohexen-1-yloxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

30 170. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethoxy)benzamide

171. 4-(difluoromethoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

172. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfanyl)benzamide

35 173. 2-[(4-chlorophenyl)sulfinyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

174. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[(4-nitrophenyl)sulfinyl]benzamide

175. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(4-methylphenyl)sulfonyl]-3-nitrobenzamide

5 176. N-(5-isopropyl-1,3-thiazol-2-yl)-3-[(trifluoromethyl)sulfonyl]benzamide

177. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxy-4-(methylsulfanyl)benzamide

178. 2-[(2-cyanophenyl)sulfonyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

10 179. N-1-,N-1--diethyl-3,6-difluoro-N-2-(5-isopropyl-1,3-thiazol-2-yl)phthalamide

180. 4-formyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

181. 2-formyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

15 182. 4-[(2,5-dimethoxyanilino)carbonyl]amino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

183. 4-(hydroxymethyl)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

184. 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)-2-nitrobenzyl acetate

20 185. 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)-2-nitrobenzyl 4-(acetylamino)-3-iodobenzoate

186. 4-(acetylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

25 187. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(2-phenylacetyl)amino]benzamide

188. 4-(acetylamino)-3-iodo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

189. 4-amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

30 190. 4-(dimethylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

191. 3-(dimethylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

192. 2-(methylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

35 193. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(trifluoromethyl)anilino]benzamide

194. 3-{{(5-bromo-1,3-dioxo-1,3-dihydro-2H-isoindol-2-  
y1)methyl}amino}-N-(5-isopropyl-1,3-thiazol-2-  
y1)benzamide

195. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(1H-pyrrol-1-  
y1)benzamide

196. 2,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-  
y1)isonicotinamide

197. 2-(4-bromophenyl)-6-(4-iodophenyl)-N-(5-isopropyl-1,3-  
thiazol-2-yl)isonicotinamide

198. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-  
(trifluoromethyl)anilino]nicotinamide

199. 2,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-  
y1)nicotinamide

200. 5,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-  
y1)nicotinamide

201. 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-6-  
methylnicotinamide

202. 2,6-dichloro-5-fluoro-N-(5-isopropyl-1,3-thiazol-2-  
y1)nicotinamide

203. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenoxyticotinamide

204. N-(5-isopropyl-1,3-thiazol-2-yl)-6-(2,2,2-  
trifluoroethoxy)nicotinamide

205. N-(5-isopropyl-1,3-thiazol-2-yl)-2,6-  
dimethoxynicotinamide

206. N-(5-isopropyl-1,3-thiazol-2-yl)-2-  
quinoxalinecarboxamide

207. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-  
pyrazinecarboxamide

208. N-(5-isopropyl-1,3-thiazol-2-yl)-8-  
quinolinecarboxamide

209. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-4-  
quinolinecarboxamide

210. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-1-phenyl-1H-  
pyrazole-4-carboxamide

211. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-1H-pyrazole-  
3-carboxamide

212. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-pyrazole-4-  
carboxamide

213. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-phenyl-2H-1,2,3-triazole-4-carboxamide

214. 2-[(2,1,3-benzoxadiazol-5-yloxy)methyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-1,3-thiazole-5-carboxamide

215. N-(5-isopropyl-1,3-thiazol-2-yl)-9H-fluorene-1-carboxamide

216. N-(5-isopropyl-1,3-thiazol-2-yl)-7-methoxy-1-benzofuran-2-carboxamide

217. N-(5-isopropyl-1,3-thiazol-2-yl)-1-[(4-methylphenyl)sulfonyl]-1H-pyrrole-3-carboxamide

218. 2-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamide

219. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamide

220. N-(5-isopropyl-1,3-thiazol-2-yl)-2-naphthamide

221. N-(5-isopropyl-1,3-thiazol-2-yl)-9,10-dioxo-9,10-dihydro-2-anthracenecarboxamide

222. N-(5-isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluorene-4-carboxamide

223. N-(5-isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluorene-1-carboxamide

224. N-(5-isopropyl-1,3-thiazol-2-yl)-8-oxo-5,6,7,8-tetrahydro-2-naphthalenecarboxamide

225. N-(5-isopropyl-1,3-thiazol-2-yl)-1,3-dioxo-1,3-dihydro-2-benzofuran-5-carboxamide

226. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-5-carboxamide

227. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-4-carboxamide

228. N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-2-phenyl-1H-indole-5-carboxamide

229. 2-butyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indole-5-carboxamide

230. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-6-carboxamide

231. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methoxy-1H-indole-2-carboxamide

232. 1-allyl-2-butyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-5-carboxamide

233. N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indole-2-carboxamide

5 234. 1-benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-1H-indole-5-carboxamide

235. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-1,2,3-benzotriazole-5-carboxamide

236. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-isoxazolecarboxamide

10 237. N-(5-isopropyl-1,3-thiazol-2-yl)-3-thiophenecarboxamide

238. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-thiophenecarboxamide

15 239. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-thiophenecarboxamide

240. 5-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-2-thiophenecarboxamide

20 241. N-(5-isopropyl-1,3-thiazol-2-yl)-3-[(2,3,3-trichloroacryloyl)amino]-2-thiophenecarboxamide

242. 5-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide

243. N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide

244. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(4-nitrophenyl)-2-furamide

25 245. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(2-nitrophenyl)-2-furamide

246. 5-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide

247. N-(5-isopropyl-1,3-thiazol-2-yl)-5-[3-(trifluoromethyl)phenyl]-2-furamide

30 248. 5-(4-chloro-2-nitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide

249. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(4-methyl-2-nitrophenyl)-2-furamide

35 250. 5-[2-chloro-5-(trifluoromethyl)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide

251. *tert*-butyl (1*R*)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethylcarbamate

252. (1R)-2-[ (5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate

253. (1S)-2-[ (5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate

5 254. (R,S)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

255. (R)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

10 256. (S)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

257. 2-(acetylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

258. (R,S)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

15 259. (R)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

260. (S)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

20 261. 3,3,3-trifluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxy-2-phenylpropanamide

262. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(1-naphthyl)acetamide

263. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-naphthyl)acetamide

25 264. 2-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

265. 2-(1,3-benzodioxol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

30 266. 2-(2,4-dinitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

267. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-methyl-1H-indol-3-yl)acetamide

268. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(1-methyl-1H-indol-3-yl)acetamide

35 269. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(5-methoxy-1H-indol-3-yl)acetamide

270. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(5-benzyloxy-1H-indol-3-yl)acetamide

271. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxy-2-methyl-1H-indol-3-yl)acetamide

272. 2-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxoacetamide

5 273. 2-(5-bromo-1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

274. 2-(5-fluoro-1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

275. 2-[1-(4-chlorobenzoyl)-5-methoxy-2-methyl-1H-indol-3-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

10 276. 3-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

277. 4-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)butanamide

15 278. N-(5-isopropyl-1,3-thiazol-2-yl)-3-(2-thienyl)propanamide

279. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-thienyl)acetamide

280. N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxo-2-(2-thienyl)acetamide

20 281. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-thienyl)acetamide

282. 2-(5-chloro-1-benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

25 283. 2-(1-benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

284. 2-[2-(formylamino)-1,3-thiazol-4-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamide

285. 2-{2-[(2-chloroacetyl)amino]-1,3-thiazol-4-yl}-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamide

30 286. 2-chloro-N-(4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}-1,3-thiazol-2-yl)acetamide

287. ethyl 2-({2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-(1H-pyrazol-3-yl)ethylidene]amino}oxy)acetate

35 288. 2-(2-furyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxoacetamide

289. 2-(5-bromo-3-pyridinyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

290. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(7-methoxy-2-oxo-2H-chromen-4-yl)acetamide

291. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenyl-3-butenamide

292. N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxo-4-(4-methyl-5-phenyl)butanamide

293. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-nitrophenyl)butanamide

294. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenylbutanamide

295. benzyl 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]-4-oxobutylcarbamate

296. methyl 5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate

297. 4-(1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)butanamide

298. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-methoxy-1-naphthyl)-4-oxobutanamide

299. 3-(2-chlorophenoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

300. 3-(4-methylphenoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

301. 3-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

302. 3-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

303. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylpentanamide

304. 3-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

305. 3-(4-methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

306. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

307. 3-phenyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

308. 2-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

309. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methylbutanamide

310. N-(5-isopropyl-1,3-thiazol-2-yl)-5-oxo-5-phenylpentanamide

311. 2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate

312. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[4-(1-oxo-1,3-dihydro-2H-isoindol-2-yl)phenyl]propanamide

313. 1-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)cyclopentanecarboxamide

5 314. 1-phenyl-N-(5-isopropyl-1,3-thiazol-2-yl)cyclopentanecarboxamide

315. 2-(3-bromo-4-methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

10 316. 2-(2-nitro-4-trifluoromethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

317. 5-cyclohexyl 1-(4-(2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl)benzyl) (2S)-2-[(tert-butoxycarbonyl)amino]pentanedioate

15 318. 2-(5,6-dimethyl-1H-benzimidazol-1-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

319. 2-[5-(4-chlorophenyl)-2H-1,2,3,4-tetraazol-2-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

320. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[5-(1-pyrrolidinyl)-2H-1,2,3,4-tetraazol-2-yl]acetamide

20 321. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methyl-1-benzothiophen-2-yl)acetamide

322. N-(5-isopropyl-1,3-thiazol-2-yl)-4,4-bis(4-methylphenyl)-3-butenamide

323. 2-cyclopropyl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

25 324. N-(4-bromo-6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexyl)benzamide

325. 2-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

30 326. benzyl 6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexylcarbamate

327. N-1-(5-isopropyl-1,3-thiazol-2-yl)-N~4--(2-propynyl)-2-butenediamide

328. 4-(2,4-dimethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide

35 329. 4-(4-benzyloxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide

330. 4-(thiophen-2-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide

331. benzyl 2-[(benzyloxy)carbonyl]amino]-5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate

5 332. 4-(1H-indol-3-yl)-N-[3-[(5-isopropyl-1,3-thiazol-2-yl)amino]-3-oxopropyl]butanamide

333. 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonylphenyl 4-chlorobenzenesulfonate

334. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(2-methoxyanilino)carbonyl]amino}benzamide

10 335. 4-[(2-(isopropylsulfonyl)acetyl]amino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

336. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(2-(phenylsulfanyl)acetyl]amino}benzamide

15 337. 4-[(diethylamino)sulfonyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

338. 2-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

339. 3,5-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

340. 3-[(2-fluoroanilino)carbonyl]amino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

20 341. N-(5-isopropyl-1,3-thiazol-2-yl)-1-phenyl-5-propyl-1H-pyrazole-4-carboxamide

342. 3-chloro-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfanyl)-2-thiophenecarboxamide

25 343. 3-iodo-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfanyl)-2-thiophenecarboxamide

344. 2-[(4-chlorophenyl)sulfonylmethyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-1,3-thiazole-5-carboxamide

30 345. 5-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(trifluoromethyl)-3-furamide

346. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,3,4,5,6-pentafluorophenyl)acetamide

35 347. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-fluorophenyl)acetamide

348. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-bromophenyl)acetamide

349. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-chlorophenyl)acetamide

350. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-nitrophenyl)acetamide

5 351. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-trifluoromethylphenyl)acetamide

352. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-methoxyphenyl)acetamide

10 353. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-dimethoxyphenyl)acetamide

354. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-difluorophenyl)acetamide

15 355. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4,5-trimethoxyphenyl)acetamide

356. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,6-dichlorophenyl)acetamide

357. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-chloro-6-fluorophenyl)acetamide

20 358. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,5-dimethoxyphenyl)acetamide

359. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,5-difluorophenyl)acetamide

360. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-bis-trifluoromethylphenyl)acetamide

25 361. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methylthiophenyl)acetamide

362. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide

30 363. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-bromophenyl)acetamide

364. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-chlorophenyl)acetamide

365. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-fluorophenyl)acetamide

35 366. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl)acetamide

367. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-trifluoromethylphenyl)acetamide

368. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methylphenyl)acetamide

369. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-dimethylaminophenyl)acetamide

5 370. 2-[1,1'-biphenyl]-4-yl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

371. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-trifluoromethylphenyl)acetamide

372. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-bromophenyl)acetamide

10 373. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-chlorophenyl)acetamide

374. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-nitrophenyl)acetamide

15 375. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide

376. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-dinitrophenyl)acetamide

377. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-20 dichlorophenyl)acetamide

378. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-difluorophenyl)acetamide

379. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-benzyloxy-3-methoxyphenyl)acetamide

25 380. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dichlorophenyl)acetamide

381. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-difluorophenyl)acetamide

382. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-30 dimethoxyphenyl)acetamide

383. 2-(2,3-dihydro-1H-inden-5-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

384. N-(5-isopropyl-1,3-thiazol-2-yl)-1-phenylcyclopropanecarboxamide

35 385. 2-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

386. 2-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

387. N-(5-isopropyl-1,3-thiazol-2-yl)-2,2-diphenylacetamide

388. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-nitrophenoxy)acetamide

5 389. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl)propanamide

390. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide

391. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-isobutylphenyl)propanamide

10 392. N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxo-2-phenylacetamide

393. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-phenylpentanamide

15 394. (E, Z)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-2-butenamide

395. N-(5-isopropyl-1,3-thiazol-2-yl)bicyclo[4.2.0]octa-1,3,5-triene-7-carboxamide

396. N-(5-isopropyl-1,3-thiazol-2-yl)-3-oxo-1-indanecarboxamide

20 397. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylbutanamide

398. tert-butyl (1S)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-1-methyl-2-oxoethylcarbamate

399. tert-butyl (1S,2S)-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl-2-methylbutylcarbamate

25 400. tert-butyl 2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

401. tert-butyl (1S)-5-amino-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonylpentylcarbamate

402. tert-butyl 4-[(imino{[(4-

30 30 methylphenyl)sulfonyl]amino}methyl)amino]-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]butylcarbamate

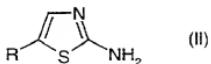
403. tert-butyl 1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl-3-(tritylamo)propylcarbamate

35 404. tert-butyl (1S)-1-(benzyloxymethyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

405. tert-butyl (1S)-1-benzyl-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

406. *tert*-butyl (1*R*)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-(benzylthiomethyl)ethylcarbamate  
 407. benzyl (3*S*)-3-[(*tert*-butoxycarbonyl)amino]-4-[(5-isopropyl-1,3-thiazol-2-yl)amino]-4-oxobutanoate  
 5 408. *tert*-butyl (2*S*)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)-1-pyrrolidinecarboxylate  
 409. *tert*-butyl (1*S*)-1-(1*H*-indol-3-ylmethyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate  
 410. *tert*-butyl (1*S*)-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)-3-(methylsulfanyl)propylcarbamate  
 10 411. *tert*-butyl (1*S*)-2-benzyloxy-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonylpropylcarbamate  
 412. *tert*-butyl (1*S*)-1-(4-benzyloxybenzyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate  
 15 413. *tert*-butyl (1*S*)-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)-2-methylpropylcarbamate  
 414. *tert*-butyl (1*S*)-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)-3-methylbutylcarbamate  
 415. benzyl (4*S*)-4-[(*tert*-butoxycarbonyl)amino]-5-oxopentanoate;  
 20 and the pharmaceutically acceptable salts thereof.

The compounds of formula (I) object of the present invention and the salts thereof can be obtained, for 25 instance, by a process comprising reacting a compound of formula (II)



with a compound of formula (III)



30 wherein R and R<sub>1</sub> are as defined above and X is hydroxy or a suitable leaving group;  
 and, if desired, converting a 2-amino-1,3-thiazole derivative of formula (I) into another such derivative of formula (I), and/or into a salt thereof.

Examples of specific compounds of formula (III) wherein X is a suitable leaving group are those wherein X represents a halogen atom, preferably chlorine or bromine.

5 It is clear to the man skilled in the art that if the compound of formula (I), prepared according to the above process is obtained as an admixture of isomers, its separation into the single isomers according to conventional techniques is still within the scope of the  
10 present invention.

Likewise, the conversion into the free compound (I) of a corresponding salt thereof, according to well-known procedures in the art, is still within the scope of the invention.

15 The above process is an analogy process which can be carried out according to well known methods.

The reaction between a compound of formula (II) and a carboxylic acid of formula (III) wherein X is a hydroxy group, can be carried out in the presence of a coupling agent or a polymer supported coupling agent such as, for instance, carbodiimide, i.e. 1,3-dicyclohexylcarbodiimide, 1,3-diisopropylcarbodiimide, 1-(3-dimethylaminopropyl)-3-ethylcarbodiimide, or N-Cyclohexylcarbodiimide N'-methylpolystyrene in a suitable solvent such as, for instance, dichloromethane, chloroform, tetrahydrofuran, diethyl ether, 1,4-dioxane, acetonitrile, toluene, or N,N-dimethylformamide at a temperature ranging from about -10°C to reflux for a suitable time, i.e. from about 30 min. to  
30 about 96 hours.

The reaction between a compound of formula (II) and a compound of formula (III) can be also carried out, for example, by a mixed anhydride method, using an alkyl chloroformate, such as ethyl, iso-butyl, or iso-propyl chloroformate, in the presence of a tertiary base, such as triethylamine, N,N-diisopropylethylamine or pyridine, in a suitable solvent such as, for instance, toluene, dichloromethane, chloroform, tetrahydrofuran, acetonitrile,

diethyl ether, 1,4-dioxane, or N,N-dimethylformamide, at a temperature ranging from about -30°C to room temperature.

The reaction between a compound of formula (II) and a carboxylic acid derivative of formula (III) wherein X is a

5 suitable leaving group can be carried out in the presence of a tertiary base, such as triethylamine, N,N-diisopropylethylamine or pyridine, in a suitable solvent, such as toluene, dichloromethane, chloroform, diethyl ether, tetrahydrofuran, acetonitrile, or N,N-  
10 dimethylformamide, at a temperature ranging from about -10°C to reflux.

Also the optional conversion of a compound of formula (I) into another compound of formula (I) can be carried out according to known methods.

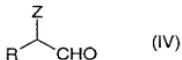
As an example, the nitro group of a compound of formula (I) may be converted into an amino group by treatment, for example, with stannous chloride in concentrated hydrochloric acid and by using, if necessary, an organic solvent such as acetic acid, 1,4-dioxane and tetrahydrofuran, at a temperature varying between room temperature and about 100°C.

Likewise, an alkylthio or an arylthio group may be converted into the corresponding alkylsulfonyl and 25 arylsulfonyl group by reaction, for example, with m-chloroperbenzoic acid in a suitable solvent such as dichloromethane or chloroform, at a temperature varying between about -5°C and room temperature.

The optional salification of a compound of formula (I) or 30 the conversion of a salt into the free compound as well as the separation of a mixture of isomers into the single isomers may be carried out by conventional methods.

The compounds of formula (II) and (III) according to the 35 process object of the present invention are known compounds or can be obtained according to known methods.

For example, a compound of formula (II) wherein R is as defined above can be obtained by reacting a compound of formula (IV)



5 wherein Z is a bromine or chlorine atom, with thiourea in a suitable solvent such as methanol, ethanol, tetrahydrofuran, 1,4-dioxane or toluene, at a temperature varying between room temperature and reflux, for a suitable time ranging from about 1 hour to about 24 hours.

10 A compound of formula (III) wherein X is a leaving group as defined above can be obtained according to conventional techniques from the corresponding carboxylic acids of formula (III) wherein X is hydroxy.

When preparing the compounds of formula (I) according to the process object of the present invention, optional functional groups within both the starting materials or the intermediates thereof, which could give rise to unwanted side reactions, need to be properly protected according to conventional techniques.

15 20 Likewise, the conversion of these latter into the free deprotected compounds may be carried out according to known procedures.

#### Pharmacology

25 The compounds of formula (I) are active as cdk/cyclin inhibitors as they gave positive results when tested according to the following procedure.

The inhibiting activity of putative cdk/cyclin inhibitors and the potency of selected compounds was determined through a method of assay based on the use of the MultiScreen-PH 96 well plate (Millipore), in which a phosphocellulose filter paper was placed at each well bottom allowing binding of positive charged substrate after 30 a washing/filtration step.

When a radioactivity labelled phosphate moiety was transferred by the ser/threo kinase to the filter-bound histone, light emitted was measured in a scintillation counter.

5 The inhibition assay of cdk2/Cyclin A activity was performed according to the following protocol:

**Kinase reaction:** 1.5  $\mu$ M histone H1 substrate, 25  $\mu$ M ATP (0.5  $\mu$ Ci  $\text{P}^{32}\gamma$ -ATP), 30 ng Cyclin A/cdk2 complex, 10  $\mu$ M 10 inhibitor in a final volume of 100  $\mu$ l buffer (TRIS HCl 10 mM pH 7.5, MgCl<sub>2</sub> 10 mM, 7.5 mM DTT) were added to each well of a 96 U bottom well plate. After 10 min at 37 °C incubation, reaction was stopped by 20  $\mu$ l EDTA 120 mM.

15 **Capture:** 100  $\mu$ l were transferred from each well to MultiScreen plate, to allow substrate binding to phosphocellulose filter. Plates were then washed 3 times with 150  $\mu$ l/well PBS Ca<sup>++</sup>/Mg<sup>++</sup> free and filtered by MultiScreen filtration system.

20 **Detection:** filters were allowed to dry at 37°C, then 100  $\mu$ l/well scintillant were added and <sup>32</sup>P labelled histone H1 was detected by radioactivity counting in the Top-Count instrument.

25 **Results:** data were analysed and expressed as % inhibition referred to total activity of enzyme (=100%).

All compounds showing inhibition  $\geq$  50 % were further analysed in order to study and define potency (IC50) as 30 well as the kinetic-profile of inhibitor through Ki calculation.

IC50 determination: the protocol used was the same described above, where inhibitors were tested at concentrations ranging from 0.0045 to 10  $\mu$ M. Experimental 35 data were analyzed by the computer program GraphPad Prizm.

Ki calculation: either the concentration of ATP and histone H1 substrate were varied: 4, 8, 12, 24, 48  $\mu$ M for ATP (containing proportionally diluted  $P^{32} \gamma$ -ATP) and 0.4, 0.8, 1.2, 2.4, 4.8  $\mu$ M for histone were used in absence and 5 presence of two different, properly chosen inhibitor concentrations.

Experimental data were analysed by the computer program SigmaPlot for  $K_i$  determination, using a random bireactant system equation:

10 
$$V_{max} \frac{(A)(B)}{aK_aK_b}$$
  
v = 
$$\frac{1 + \frac{(A)}{K_a} + \frac{(B)}{K_b} + \frac{(A)(B)}{aK_aK_b}}{1 + \frac{(A)}{K_a} + \frac{(B)}{K_b}}$$

15 where A=ATP and B=histone H1.

Following the method above described, a representative compound of formula (I) of the invention, which is 2-[4-(dimethylamino)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide, showed an inhibiting activity towards the 20 cdk2/cyclin A complex corresponding to 0.1  $\mu$ M ( $K_i$ ).

In addition, the inhibiting activity of putative cdk/cyclin 25 inhibitors and the potency of selected compounds was determined through a method of assay based on the use of a SPA (Scintillation Proximity Assay) 96 well plate assay. The assay is based on the ability of streptavidin coated SPA beads to capture a biotinylated peptide derived from a phosphorylation site of histone.

30 When a radioactivity labelled phosphate moiety was transferred by the ser/threo kinase to the biotinylated histone peptide, light emitted was measured in a scintillation counter.

35 The inhibition assay of cdk5/p25 activity was performed according to the following protocol:

**Kinase reaction:** 1.0  $\mu$ M biotinylated histone peptide substrate, 0.25 uCi P33g-ATP, 4 nM cdk2/p25 complex, 0-100  $\mu$ M inhibitor in a final volume of 100  $\mu$ l buffer (Hepes 20 mM pH 7.5, MgCl<sub>2</sub> 15 mM, 1 mM DTT) were added to each well 5 of a 96 U bottom well plate. After 20 min at 37 °C incubation, the reaction was stopped by the addition of 500 ug SPA beads in phosphate-buffered saline containing 0.1% Triton X-100, 50  $\mu$ M ATP and 5 mM EDTA. The beads were allowed to settle, and the radioactivity incorporated in 10 the 33P-labelled peptide was detected in a Top Count scintillation counter.

**Results:** Data were analyzed and expressed as % Inhibition using the formula:

15  $100 \times (1 - (\text{Unknown} - \text{Bkgd}) / (\text{Enz. Control} - \text{Bkgd}))$

IC50 values were calculated using a variation of the four parameter logistics equation:

$$Y = 100 / [1 + 10^{((\text{LogEC50} - X) * \text{Slope})}]$$

20

Where X = log(uM) and Y = % Inhibition.

The compounds of formula (I) are therefore useful to restrict the unregulated proliferation of tumor cells, 25 hence in therapy in the treatment of various tumors such as, for instance, carcinomas, e.g. mammary carcinoma, lung carcinoma, bladder carcinoma, colon carcinoma, ovary and endometrial tumors, sarcomas, e.g. soft tissue and bone sarcomas, and the hematological malignancies such as, e.g., 30 leukemias.

In addition, the compounds of formula (I) are also useful in the treatment of other cell proliferative disorders such as psoriasis, vascular smooth cell proliferation associated with atherosclerosis and post-surgical stenosis and 35 restenosis and in the treatment of Alzheimer's disease.

The compounds of the present invention can be administered either as single agents or, alternatively, in combination with known anticancer treatments such as radiation therapy or chemotherapy regimen in combination with cytostatic or 5 cytotoxic agents.

As an example, the above compounds can be administered in combination with one or more chemotherapeutic agents such as, for instance, taxane, taxane derivatives, CPT-11, camptothecin derivatives, anthracycline glycosides, e.g. 10 doxorubicin or epirubicin, etoposide, navelbine, vinblastine, carboplatin, cisplatin and the like, optionally within liposomal formulations thereof.

15 The compounds of formula (I) of the present invention, suitable for administration to a mammal, e.g. to humans, can be administered by the usual routes and the dosage level depends upon the age, weight, conditions of the patient and the administration route.

20 For example, a suitable dosage adopted for oral administration of a compound of formula (I) may range from about 10 to about 500 mg pro dose, from 1 to 5 times daily. The compounds of the invention can be administered in a variety of dosage forms, e.g. orally, in the form of tablets, capsules, sugar or film coated tablets, liquid 25 solutions or suspensions; rectally in the form of suppositories; parenterally, e.g. intramuscularly, or by intravenous and/or intrathecal and/or intraspinal injection or infusion.

30 The present invention also includes pharmaceutical compositions comprising a compound of formula (I), or a pharmaceutically acceptable salt thereof, in association with a pharmaceutically acceptable excipient (which can be a carrier or a diluent).

35 The pharmaceutical compositions containing the compounds of the invention are usually prepared following conventional methods and are administered in a pharmaceutically suitable form.

For example, the solid oral forms may contain, together with the active compound, diluents, e.g. lactose, dextrose, saccharose, sucrose, cellulose, corn starch or potato starch; lubricants, e.g. silica, talc, stearic acid, 5 magnesium or calcium stearate, and/or polyethylene glycols; binding agents, e.g. starches, arabic gum, gelatine, methylcellulose, carboxymethylcellulose or polyvinyl pyrrolidone; disaggregating agents, e.g. a starch, alginic acid, alginates or sodium starch glycolate; effervescent 10 mixtures; dyestuffs; sweeteners; wetting agents such as lecithin, polysorbates, laurylsulphates; and, in general, non-toxic and pharmacologically inactive substances used in pharmaceutical 15 preparations may be manufactured in known manner, for example, by means of mixing, granulating, tabletting, sugar-coating, or film-coating processes.

The liquid dispersions for oral administration may be e.g. syrups, emulsions and suspensions.

The syrups may contain as carrier, for example, saccharose or saccharose with glycerine and/or mannitol and/or sorbitol.

The suspensions and the emulsions may contain as carrier, for example, a natural gum, agar, sodium alginate, pectin, methylcellulose, carboxymethylcellulose, or polyvinyl 25 alcohol.

The suspension or solutions for intramuscular injections may contain, together with the active compound, a pharmaceutically acceptable carrier, e.g. sterile water, olive oil, ethyl oleate, glycols, e.g. propylene glycol, 30 and, if desired, a suitable amount of lidocaine hydrochloride. The solutions for intravenous injections or infusions may contain as carrier, for example, sterile water or preferably they may be in the form of sterile, aqueous, isotonic saline solutions or they may contain as a carrier propylene glycol.

The suppositories may contain together with the active compound a pharmaceutically acceptable carrier, e.g. cocoa

butter, polyethylene glycol, a polyoxyethylene sorbitan fatty acid ester surfactant or lecithin.

5 The following examples illustrate but do not limit the present invention.

**Example 1**

**Preparation of Ethyl 3-[(5-bromo-1,3-thiazol-2-yl)amino]-3-oxopropanoate**

10 Ethyl malonyl chloride (0.88 ml; 6.99 mmol) was added to a mixture of 2-amino-5-bromothiazole hydrobromide (1.30 g; 5.00 mmol) and Et3N (2.08 ml; 14.94 mmol) in THF (6 ml) at 0-5°C. The mixture was stirred at room temperature overnight, then the reaction was quenched with potassium 15 sarcosinate (0.25 g; 2.00 mmol) and water (12 ml). The product was isolated by filtration as a white solid (0.75 g, 51%): m.p. 165-166°C.

20  $^1\text{H-NMR}$  (CDCl<sub>3</sub>)  $\delta$  ppm: 10.80 (bs, 1H, CONH); 7.38 (s, 1H, thiazole CH); 4.28 (q, J = 7.3 Hz, 2H, COOCH<sub>2</sub>CH<sub>3</sub>); 3.56 (s, 2H, COCH<sub>2</sub>CO); 1.32 (t, J = 7.3 Hz, 2H, COOCH<sub>2</sub>CH<sub>3</sub>).

Analogously, the following products can be prepared:

N-(5-bromo-1,3-thiazol-2-yl)-2-phenyl-acetamide  
m.p. 206-207°C

25  $^1\text{H-NMR}$  (DMSO-d<sub>6</sub>)  $\delta$  ppm: 3.76 (s, 2H, COCH<sub>2</sub>Ph); 7.2-7.3 (m, 5H, Ph); 7.54 (s, 1H, thiazole CH); 12.80 (bs, 1H, CONH); N-(5-bromo-1,3-thiazol-2-yl)-benzamide  
m.p. 126-128°C

30  $^1\text{H-NMR}$  (DMSO-d<sub>6</sub>)  $\delta$  ppm: 12.90 (bs, 1H, CONH); 8.07, 7.93 (m, 2H, o-Ph hydrogens); 7.63 (s, 1H, thiazole CH); 7.62, 7.53, 7.48 (m, 3H, m- and p-Ph hydrogens);

Ethyl 4-[(5-bromo-1,3-thiazol-2-yl)amino]-4-oxobutanoate.

**Example 2**

**Preparation of N-(5-Bromo-thiazol-2-yl)-3-hydroxy-propionamide**

35

A mixture of LiBH<sub>4</sub> (44 mg, 2.02 mmol), ethyl 3-[(5-bromo-1,3-thiazol-2-yl)amino]-3-oxopropanoate (340 mg, 1.16 mmol), methanol (0.082 ml, 2.02 mmol), and Et<sub>2</sub>O (50 ml) was refluxed for 20 min. The reaction was quenched with 1 N hydrochloric acid with ice-cooling.

The mixture was then diluted with water and extracted with dichloromethane. The extract was dried and the solvent was evaporated under reduced pressure. Purification by silica gel chromatography (dichloromethane/methanol=98:2 and then 95:5) yielded the title compound as a white solid (0.17 g; 52%).

m.p. 182-184°C (dec.)

<sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ ppm: 10.20 (bs, 1H, CONH); 7.35 (s, 1H, thiazole CH); 4.04 (t, J = 5.4 Hz, 2H, COCH<sub>2</sub>CH<sub>2</sub>OH); 2.74 (t, J = 5.4 Hz, 2H, COCH<sub>2</sub>CH<sub>2</sub>OH).

Analogously, starting from the corresponding ester derivative the following product can be prepared:  
N-(5-Bromo-1,3-thiazol-2-yl)-4-hydroxybutanamide.

20

### Example 3

#### Preparation of N-(5-Bromo-thiazol-2-yl)-2-ethoxy-acetamide

EDCI (0.53 g, 2.78 mmol) was added to a solution of ethoxyacetic acid (0.29 g, 2.78 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (5 ml) under ice-cooling.

After stirring for 1 h, a solution of 2-amino-5-bromothiazole hydrobromide (0.60 g, 2.31 mmol) and diisopropylethylamine (0.40 ml, 2.34 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (5 ml) was added dropwise, and the entire mixture was kept at 0°C for 1 h, then at room temperature overnight.

The solution was evaporated and the residue partitioned between ethyl acetate and water. The ethyl acetate layer was further washed with water, 5% citric acid, water, saturated sodium bicarbonate, and water.

35 Drying over sodium sulfate and evaporation gave a solid which was triturated with isopropyl ether to give the title compound as a beige solid (0.43 g; 70%)

m.p. 100-102°C

<sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ ppm: 9.64 (bs, 1H, CONH); 7.38 (s, 1H, thiazole CH); 4.16 (s, 2H, COCH<sub>2</sub>O); 3.65 (q, J = 6.8 Hz, 2H, OCH<sub>2</sub>CH<sub>3</sub>); 1.29 (t, J = 6.8 Hz, 3H, OCH<sub>2</sub>CH<sub>3</sub>).

5 Analogously, the following products can be prepared:

tert-butyl 3-[(5-bromo-1,3-thiazol-2-yl)amino]-3-oxopropylcarbamate;

Benzyl 4-[(5-bromo-1,3-thiazol-2-yl)amino]-4-oxobutylcarbamate;

10 tert-butyl 4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}phenylcarbamate

tert-butyl 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonylphenylcarbamate

15 tert-butyl 4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}-1,3-thiazol-2-ylcarbamate;

N-(5-bromo-1,3-thiazol-2-yl)-2-bromoacetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-bromoacetamide;

2-N-[2-(3-pyridyl)-acetyl-amino]-5-bromo-thiazole

m.p. 232-235°C

20 <sup>1</sup>H-NMR (DMSO-d<sub>6</sub>): 3.82 (s, 2H, COCH<sub>2</sub>Ph); 7.34 (dd, J=4.4, 7.7 Hz, 1H, H<sub>5</sub> Py); 7.55 (s, 1H, thiazole CH); 7.71 (ddd, J=1.6, 2.2, 7.7 Hz, 1H, H<sub>4</sub> Py); 8.45 (dd, J=1.6, 4.9 Hz, 1H, H<sub>6</sub> Py); 8.49 (d, J=2.2 Hz, 1H, H<sub>2</sub> Py); 12.65 (s, 1H, CONH);

25 2-N-[2-(3-pyridyl)-acetyl-amino]-5-isopropyl-thiazole  
m.p. 178-180°C (dec.)

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.20 (bs, 1H, CONH); 8.45, 7.7, 7.35 (m, 4H, Py); 7.17 (s, 1H, thiazole CH); 3.78 (s, 2H, COCH<sub>2</sub>); 3.14 (m, 1H, CH(Me)<sub>2</sub>); 1.22 (d, 6H, CHMe<sub>2</sub>);

30 N-(5-bromo-1,3-thiazol-2-yl)-2-(3-hydroxyphenyl)acetamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-hydroxyphenyl)acetamide  
m.p. 206-208°C.

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.1 (bs, 1H, CONH); 9.34 (s, 1H,

35 OH); 7.14 (s, 1H, thiazole CH); 7.1 (t, 1H, H<sub>5</sub> Ph); 6.6-6.7

(m, 3H, H<sub>2</sub>, H<sub>4</sub>, H<sub>6</sub> Ph); 3.6 (s, 2H, COCH<sub>2</sub>); 3.08 (ept, 1H, CHMe<sub>2</sub>); 1.22 (d, 6H, CHMe<sub>2</sub>);

N-(5-bromo-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-

5 methoxyphenyl)acetamide

m.p. 97-98°C.

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.12 (s, 1H, CONH); 7.21 (dd, 1H, H<sub>5</sub>

Ph); 7.14 (d, 1H, thiazole CH); 6.87 (m, 2H, H<sub>2</sub>, H<sub>6</sub> Ph);

6.81 (ddd, 1H, H<sub>4</sub> Ph); 3.72 (s, 3H, OMe); 3.67 (s, 2H,

10 COCH<sub>2</sub>); 3.07 (m, 1H, CHMe<sub>2</sub>); 1.22 (d, 6H, CHMe<sub>2</sub>);

N-(5-bromo-1,3-thiazol-2-yl)-2-(3-chorophenyl)acetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-chorophenyl)acetamide

m.p. 116-118°C.

<sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ ppm: 11.8 (bs, 1H, CONH); 7.32 (s, 1H, H<sub>2</sub>

15 Ph); 7.24 (m, 3H, H<sub>4</sub>, H<sub>5</sub>, H<sub>6</sub> Ph); 7.04 (s, 1H, thiazole CH); 3.76 (s, 2H, COCH<sub>2</sub>); 3.13 (m, 1H, CHMe<sub>2</sub>); 1.31 (d, 6H, CHMe<sub>2</sub>);

N-(5-bromo-1,3-thiazol-2-yl)-2-(4-hydroxyphenyl)acetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-

20 hydroxyphenyl)acetamide

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.07 (bs, 1H, CONH); 9.33 (sb, 1H, OH);

7.17-6.7 (m, 5H, Ar+ CHthiazole); 3.60 (s, 2H, COCH<sub>2</sub>);

3.1 (m, 1H, CHMe<sub>2</sub>); 1.23 (d, 6H, CHMe<sub>2</sub>);

N-(5-bromo-1,3-thiazol-2-yl)-2-(3,4-

25 dihydroxyphenyl)acetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-

dihydroxyphenyl)acetamide

m.p. 168-169°C.

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.01 (bs, 1H, CONH); 8.79 (sb, 2H, 2

30 OH); 7.12 (s, 1H, thiazole CH); 6.69 (d, 1H, H<sub>2</sub> Ph); 6.63

(d, 1H, H<sub>5</sub> Ph); 6.52 (dd, 1H, H<sub>6</sub> Ph); 3.48 (s, 2H, COCH<sub>2</sub>);

3.06 (m, 1H, CHMe<sub>2</sub>); 1.22 (d, 6H, CHMe<sub>2</sub>);

N-(5-bromo-1,3-thiazol-2-yl)-2-(4-hydroxy-3-

methoxyphenyl)acetamide;

35 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxy-3-

methoxyphenyl)acetamide

m.p. 115-116°C.

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.0 (bs, 1H, CONH); 8.80 (s, 1H, OH); 7.12 (d, 1H, thiazole CH); 6.88 (s, 1H, H2 Ph); 6.68 (m, 2H, H5, H6 Ph); 3.73 (s, 3H, OMe); 3.56 (s, 2H, COCH<sub>2</sub>); 3.07 (m, 1H, CHMe<sub>2</sub>); 1.22 (d, 6H, CHMe<sub>2</sub>);

5 N-(5-bromo-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide  
m.p. 129-130°C.

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.08 (s, 1H, CONH); 7.21 (dd, 2H, H2, H6 Ph); 7.13 (d, 1H, thiazole CH); 6.87 (dd, 2H, H3, H5 Ph); 3.70 (s, 3H, OMe); 3.62 (s, 2H, COCH<sub>2</sub>); 3.06 (m, 1H, CHMe<sub>2</sub>); 1.22 (d, 6H, CHMe<sub>2</sub>);  
N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-acetamide  
m.p. 135-137°C

15 <sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.20 (bs, 1H, CONH); 7.29 (m, 5H, Ph); 7.13 (s, 1H, thiazole CH); 3.70 (s, 2H, COCH<sub>2</sub>); 3.07 (m, 1H, CHMe<sub>2</sub>); 1.22 (d, 6H, CHMe<sub>2</sub>);  
2-[3-(3-chloropropoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

20 m.p. 91-92°C

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.08 (bs, 1H, CONH); 7.21 (t, 1H, H5 Ph); 7.13 (s, 1H, thiazole CH); 6.8-6.9 (m, 3H, H2, H4, H6 Ph); 4.05 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Cl); 3.77 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Cl); 3.67 (s, 2H, COCH<sub>2</sub>); 3.07 (ept, 1H, CHMe<sub>2</sub>); 2.14 (quint, 2H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Cl); 1.22 (d, 6H, CHMe<sub>2</sub>); and  
2-[3-(2-chloroethoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
m.p. 134-135°C

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.09 (bs, 1H, CONH); 7.22 (t, 1H, H5 Ph); 7.13 (s, 1H, thiazole CH); 6.8-6.9 (m, 3H, H2, H4, H6 Ph); 4.2 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>Cl); 3.91 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>Cl); 3.67 (s, 2H, COCH<sub>2</sub>); 3.07 (ept, 1H, CHMe<sub>2</sub>); 1.22 (d, 6H, CHMe<sub>2</sub>).

#### Example 4

35 Preparation of N-(5-bromo-thiazol-2-yl)-4-sulfamoyl-benzamide.

To a mixture of 4-sulfamoylbenzoic acid (1.0 g, 4.97 mmol), Et3N (1.5 ml, 10.78 mmol), DMF (5 ml) and THF (5 ml) isobutyl chloroformate (0.70 ml, 5.36 mmol) was added dropwise at -10°C.

5 After stirring for 1 h, a solution of 2-amino-5-bromothiazole hydrobromide (1.55 g, 5.96 mmol) and Et3N (0.83 ml, 5.96 mmol) in DMF (6 ml) and THF (4 ml) was added dropwise to the mixture at the same temperature.

10 The resulting mixture was gradually warmed to room temperature over a period of 3 h and then concentrated by evaporation of the solvent in vacuo. To the resultant residue AcOEt and 5% aqueous NaHCO3 were added. The separated organic phase was washed with water, dried over anhydrous Na2SO4, and concentrated under reduced pressure.

15 The residual solid was purified by flash chromatography (dichloromethane/methanol/30% aqueous ammonia=95:5:0.5) to afford the title compound as a yellow solid (0.77 g, 43%) m.p. 268-270°C.

20 <sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 7.54 (s, 2H, SO<sub>2</sub>NH<sub>2</sub>); 7.67 (s, 1H, thiazole CH); 7.94 (d, J=8.8 Hz, 2H, H<sub>3</sub> and H<sub>5</sub> Ph); 8.21 (d, J=8.8 Hz, 2H, H<sub>2</sub> and H<sub>6</sub> Ph); 13.10 (bs, 1H, CONH).

Analogously, the following product can be prepared:

N-(5-isopropyl-thiazol-2-yl)-4-sulfamoyl-benzamide  
m.p. 222-224°C.

25 <sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.65 (bs, 1H, CONH); 8.18 (dd, 2H, H<sub>2</sub>, H<sub>6</sub> Ph); 7.92 (dd, 2H, H<sub>3</sub>, H<sub>5</sub> Ph); 7.51 (s, 2H, SO<sub>2</sub>NH<sub>2</sub>); 7.25 (s, 1H, thiazole CH); 3.13 (m, 1H, CH<sub>2</sub>Me<sub>2</sub>); 1.28 (d, 6H, CH<sub>2</sub>Me<sub>2</sub>).

30 **Example 5**

Preparation of 4-amino-N-(5-bromo-1,3-thiazol-2-yl)butanamide hydrobromide

A solution (1.3 ml) of hydrogen bromide in glacial acetic acid (33%) was added to benzyl 4-[(5-bromo-1,3-thiazol-2-yl)amino]-3-oxobutylcarbamate (0.72 g, 1.81 mmol) and the mixture was stirred at room temperature for 1 h.

Ether was added and the solid was filtered and washed with ether. The crude product was recrystallized from MeOH/ether to afford the title compound as a beige solid (0.38 g, 61%), m.p. 211-213°C (dec.).

5  $^1\text{H-NMR}$  (DMSO- $d_6$ )  $\delta$  ppm: 1.84 (m, 2H,  $\text{COCH}_2\text{CH}_2\text{CH}_2\text{NH}_2$ ); 2.53 (t,  $J=6.8$  Hz, 2H,  $\text{COCH}_2\text{CH}_2\text{CH}_2\text{NH}_2$ ); 2.81 (m, 2H,  $\text{COCH}_2\text{CH}_2\text{CH}_2\text{NH}_2$ ); 7.68 (bs, 3H,  $\text{NH}_3^+$ ); 12.42 (s, 1H, CONH).

**Example 6**

10 Preparation of 3-amino-N-(5-bromo-1,3-thiazol-2-yl)propionamide hydrochloride

A solution 3.6 N HCl in isopropanol (14 ml) was added to tert-butyl 3-[(5-bromo-1,3-thiazol-2-yl)amino]-4-oxopropylcarbamte (0.90 g, 2.57 mmol) and the mixture was stirred at room temperature overnight. The solvent was evaporated and the residual solid was triturated in ether, filtered and dried in vacuo to afford the title compound as a white solid (0.73 g, quantitative yield) m.p. 255°C ca. (dec.).

20  $^1\text{H-NMR}$  (DMSO- $d_6$ )  $\delta$  ppm: 2.83 (t,  $J=6.8$  Hz, 2H,  $\text{COCH}_2\text{CH}_2\text{NH}_2$ ); 3.07 (q,  $J=6.4$  Hz, 2H,  $\text{COCH}_2\text{CH}_2\text{NH}_2$ ); 7.55 (s, 1H, thiazole CH); 7.96 (bs, 3H,  $\text{NH}_3^+$ ); 12.58 (s, 1H, CONH).

Analogously, the following compounds can be prepared:

25 2-(4-aminophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
 $^1\text{H-NMR}$  (DMSO- $d_6$ )  $\delta$  ppm: 1.22 (d, 6H,  $\text{CHMe}_2$ ); 3.07 (m, 1H,  $\text{CHMe}_2$ ); 3.47 (s, 2H,  $\text{COCH}_2$ ); 4.94 (s, 2H,  $\text{NH}_2$ ); 6.48 (m, 2H, H3, H5 Ph); 6.93 (m, 2H, H2, H6 Ph); 7.12 (d, 1H, CH thiazole); 12.00 (s, 1H, CONH).

30 4-amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
 $^1\text{H-NMR}$  (DMSO- $d_6$ )  $\delta$  ppm: 1.29 (d, 6H,  $\text{CHMe}_2$ ); 3.12 (m, 1H,  $\text{CHMe}_2$ ); 6.58 (m, 2H, H3, H5 Ph); 7.18 (d, 1H, CH thiazole); 7.82 (m, 2H, H2, H6 Ph); 12.80 (bs, 1H, CONH).

35 2-(2-amino-1,3-thiazol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
m.p. 204-206°C ca. (dec.)

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 1.24 (d, 6H, CHMe<sub>2</sub>); 3.10 (m, 1H, CHMe<sub>2</sub>); 3.54 (s, 2H, COCH<sub>2</sub>); 6.30 (s, 1H, H5 thiazole); 6.88 (s, 2H, NH<sub>2</sub>); 7.13 (s, 1H, H4 thiazole); 11.90 (s, 1H, CONH).

5

**Example 7****Preparation of N-(5-isopropyl-1,3-thiazol-2-yl)-butanamide**

Triethylamine (0.97 ml; 6.34 mmol) and butanoyl chloride (0.52 ml; 5.07 mmol) were added in this order to a solution 10 of 2-amino-5-isopropyl-1,3-thiazole (0.6 g; 4.23 mmol) in dichloromethane (8 ml), cooled to -5°C.

The reaction mixture was stirred at -5°C for 2 hours and then warmed to room temperature. After additional 4 hours, 15 the organic layer was washed with water, saturated sodium bicarbonate, 1N hydrochloric acid, brine, dried over sodium sulfate and evaporated. The residue was recrystallized from cyclohexane to yield 0.45 g (50%) of the title compound as a colourless solid (m.p. 95-97°C)

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 11.82 (s, 1H, CONH); 7.11 (s, 1H, thiazole CH); 3.08 (m, 1H, CHMe<sub>2</sub>); 2.34 (t, J = 7.1 Hz, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 1.58 (m, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 1.23 (d, J = 6.6 Hz, 6H, (CH<sub>3</sub>)<sub>2</sub>CH); 0.87 (t, J = 7.1 Hz, 3H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>).

Analogously, the following compounds can be prepared:

25 N-(5-bromo-1,3-thiazol-2-yl)-butanamide  
m.p. 163-164°C

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.27 (bs, 1H, CONH); 7.50 (s, 1H, thiazole CH); 2.39 (t, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 1.59 (m, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 0.87 (t, 3H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>);  
30 N-(5-chloro-1,3-thiazol-2-yl)-butanamide  
m.p. 170-171°C

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.25 (bs, 1H, CONH); 7.46 (s, 1H, thiazole CH); 2.38 (t, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 1.59 (m, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 0.87 (t, 3H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>);  
35 N-(5-phenyl-1,3-thiazol-2-yl)-butanamide  
m.p. 183-184°C

5       $^1\text{H-NMR}$  (DMSO-d<sub>6</sub>)  $\delta$  ppm: 12.13 (s, 1H, CONH), 7.84 (s, 1H, thiazole CH); 7.58 (d,  $J$  = 6.8 Hz, 2H, o-Ph hydrogens); 7.39 (dd,  $J$  = 6.8 and 7.8 Hz, 2H, m-Ph hydrogens); 7.28 (t,  $J$  = 7.8 Hz, 1H, p-Ph hydrogens); 2.41 (t,  $J$  = 7.3 Hz, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 1.61 (m, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 0.89 (t,  $J$  = 7.3 Hz, 3H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>);  
N-(5-nitro-1,3-thiazol-2-yl)-butanamide  
m.p. 175-176°C

10      $^1\text{H-NMR}$  (DMSO-d<sub>6</sub>)  $\delta$  ppm: 13.02 (s, 1H, CONH); 8.60 (s, 1H, thiazole CH); 2.48 (t,  $J$  = 7.3 Hz, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 1.62 (m, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 0.89 (t,  $J$  = 7.3 Hz, 3H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>);  
N-(5-methyl-1,3-thiazol-2-yl)-butanamide  
m.p. 137-138°C

15      $^1\text{H-NMR}$  (CDCl<sub>3</sub>)  $\delta$  ppm: 11.89 (s, 1H, CONH); 7.04 (s, 1H, thiazole CH); 2.48 (t,  $J$  = 7.3 Hz, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 2.41 (s, 3H, CH<sub>3</sub>); 1.80 (m, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 1.02 (t,  $J$  = 7.3 Hz, 3H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>);  
N-(5-benzyl-1,3-thiazol-2-yl)-butanamide  
20     m.p. 147-149°C

25      $^1\text{H-NMR}$  (CDCl<sub>3</sub>)  $\delta$  ppm: 7.23 (m, 5H, Ph); 7.07 (s, 1H, thiazole CH); 4.08 (s, 2H, CH<sub>2</sub>Ph); 2.45 (t,  $J$  = 7.8 Hz, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 1.76 (m, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 0.97 (t,  $J$  = 7.8 Hz, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>);  
N-(5-isobutyl-1,3-thiazol-2-yl)-butanamide  
m.p. 58-60°C

30      $^1\text{H-NMR}$  (CDCl<sub>3</sub>)  $\delta$  ppm: 7.03 (s, 1H, thiazole CH); 2.61 (d,  $J$  = 7.3 Hz, 2H, Me<sub>2</sub>CHCH<sub>2</sub>); 2.45 (t,  $J$  = 7.8 Hz, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 1.81 (m, 1H, Me<sub>2</sub>CHCH<sub>2</sub>); 1.78 (m, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 1.01 (t,  $J$  = 7.8 Hz, 3H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 0.95, 0.93 (s, 6H, Me<sub>2</sub>CHCH<sub>2</sub>);  
N-(5-cyclopropyl-1,3-thiazol-2-yl)-butanamide;

35     N-{5-[2-(methylsulfonyl)ethyl]-1,3-thiazol-2-yl}-butanamide  
m.p. 153-155°C

40      $^1\text{H-NMR}$  (CDCl<sub>3</sub>)  $\delta$  ppm: 11.01 (s, 1H, CONH); 7.21 (s, 1H, thiazole CH); 3.34 (m, 4H, CH<sub>3</sub>SO<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>); 2.90 (s, 3H,

CH<sub>3</sub>SO<sub>2</sub>); 2.48 (t, J = 7.3 Hz, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 1.80 (m, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 1.02 (t, J = 7.3 Hz, 3H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>);

N-[5-(2-methylthioethyl)-1,3-thiazol-2-yl]-butanamide

m.p. 67-69°C

5 <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ ppm: 11.63 (bs, 1H, NHCO), 7.26 (s, 1H, thiazole CH), 3.06 (t, J = 7.0 Hz, 2H, CH<sub>3</sub>SCH<sub>2</sub>CH<sub>2</sub>); 2.77 (t, J = 7.0 Hz, 2H, CH<sub>3</sub>SCH<sub>2</sub>CH<sub>2</sub>); 2.48 (t, J = 7.3 Hz, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 2.14 (s, 3H, CH<sub>3</sub>S); 1.80 (m, 2H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 1.02 (t, J = 7.3 Hz, 3H, COCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>);

10 N-[5-[2-(methoxycarbonyl)ethyl]-1,3-thiazol-2-yl]-butanamide;

N-[5-(3-methoxy-propyl)-1,3-thiazol-2-yl]-butanamide  
m.p. 80-82°C

15 <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ ppm: 11 (sb, 1H, NHCO); 7.07 (s, 1H, H<sub>4</sub> thiazole); 3.41 (t, 2H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>OMe); 3.34 (s 3H, OCH<sub>3</sub>); 2.85 (t, 2H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>OMe); 2.46 (t, 2H, NHCOCH<sub>2</sub>); 1.91 (m, 2H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>OMe); 1.80 (t, 2H, NHCOCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>); 1.01 (t, 3H, NHCOCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>);  
N-[5-(2-ethoxy-ethyl)-1,3-thiazol-2-yl]-butanamide

20 m.p. 74-76°C

25 <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ ppm: 7.14 (s, 1H, H<sub>4</sub> thiazole); 3.64 (t, 2H, CH<sub>2</sub>CH<sub>2</sub>OEt); 3.52 (q 2H, OCH<sub>2</sub>CH<sub>3</sub>); 3.02 (t, 2H, CH<sub>2</sub>CH<sub>2</sub>OEt); 2.47 (t, 2H, NHCOCH<sub>2</sub>); 1.80 (m, 2H, NHCOCH<sub>2</sub>CH<sub>2</sub>); 1.23 (t, 3H, OCH<sub>2</sub>CH<sub>3</sub>); 1.02 (t, 3H, NHCOCH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>);

30 N-[5-(indol-3-yl-methyl)-1,3-thiazol-2-yl]-butanamide  
m.p. 240-242°C

35 <sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ ppm: 9.95 (bs, 1H, CONH); 8.00 (bs, 1H, NH indole); 7.54 (d, 1H, H<sub>4</sub> indole); 7.36 (d, 1H, H<sub>7</sub> indole); 7.19 (m, 1H, H<sub>6</sub> indole); 7.17 (s, 1H, thiazole CH); 7.09 (m, 1H, H<sub>5</sub> indole); 7.07 (s, 1H, H<sub>2</sub> indole); 4.23 (s, 2H, CH<sub>2</sub>); 2.39 (m 2H, CH<sub>2</sub>CO); 1.73 (m, 2H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>); 0.96 (t, 3H, CH<sub>3</sub>CH<sub>2</sub>CH<sub>3</sub>).

N-[5-(3-dimethylaminoimino-butyl)-1,3-thiazol-2-yl]-butanamide.

**2-amino-5-isopropyl-1,3-thiazole**

2 ml (18.6 mmol) of 3-methylbutyraldehyde were dissolved in 15 ml of dioxane. 40.4 ml (18.6 mmol) of a solution 2 % v/v of bromine in dioxane was dropped therein at 0°C.

5 The mixture was maintained at room temperature under stirring for 2 hours, then 2.83 g (37.2 mmol) of thiourea and 5 ml of ethanol were added.

After 6 hours at room temperature the solution was evaporated to dryness, the residue was dissolved in

10 methylene chloride and the product extracted with 1M hydrochloric acid; the aqueous layer was made basic by using 30% ammonium hydrate and extracted again with methylene chloride. The organic phase was dried over sodium sulfate and evaporated under vacuum. The residue was

15 chromatographed on a silica gel column, eluting with cyclohexane-ethylacetate to give 1.1 g (42% yield) of the title compound.

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 6.6 (s, 2H, NH<sub>2</sub>); 6.58 (s, 1H, thiazole CH); 2.9 (m, 1H, CHMe<sub>2</sub>); 1.18 (s, 3H, MeCHMe); 20 1.17 (s, 3H, MeCHMe).

Analogously the following products can be prepared starting from the suitable aldehyde:

**2-amino-5-isobutyl-1,3-thiazole**

25 <sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 6.61 (sb, 2H, NH<sub>2</sub>); 6.56 (s, 1H, thiazole CH); 2.39 (dd, 2H, CH<sub>2</sub>CHMe<sub>2</sub>); 1.65 (m, 1H, CHMe<sub>2</sub>); 0.85 (d, 6H, CHMe<sub>2</sub>);

2-amino-5-phenyl-1,3-thiazole;

2-amino-5-benzyl-1,3-thiazole;

30 <sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 7.3-7.2 (m, 5H, Ph); 6.68 (s, 1H, thiazole CH); 6.67 (sb, 2H, NH<sub>2</sub>); 3.87 (s, 2H, CH<sub>2</sub>Ph);

2-amino-5-(3-indolylmethyl)-1,3-thiazole;

2-amino-5-ethoxyethyl-1,3-thiazole;

2-amino-5-methoxypropyl-1,3-thiazole;

35 2-amino-5-cyclopropyl-1,3-thiazole;

2-amino-5-methylthioethyl-1,3-thiazole;

2-amino-5-formyl-1,3-thiazole;

2-amino-5-(3-dimethylaminoimino)butyl-1,3-thiazole.

**Example 9**

**4-ethoxy-1-butanol**

5 85 mg (0.004 mmol) of sodium were dissolved in 50 ml of methanol and 8.7 g (0.23 mol) of sodium borohydride were added. A solution of 4.6 g (0.032 mol) of methyl 4-ethoxybutanoate in 20 ml of methanol was dropped to the mixture under stirring. The reaction is maintained at reflux for 6  
10 hours, then 300 ml of brine were added and the product was extracted with methylene chloride. The organic layer was dried over anhydrous sodium sulfate and evaporated to dryness to give 2.25 g (61% yield) of the title compound.

15 Analogously the following products can be prepared starting from the suitable ester:  
2-cyclopropyl-1-ethanol;  
3-(3-indolyl)-1-propanol; and  
5-dimethylaminoimino-1-hexanol.

20

**Example 10**

**Methyl 3-(3-indolyl)-propanoate**

25 2 g (10.57 mmol) of 3-indolepropionic acid were dissolved in 50 ml of methanol. The solution was cooled to 0°C and 5 ml of sulfuric acid 96% were dropped under stirring. The solution was maintained at room temperature overnight and then poured onto ice-water, basified with 30 % ammonium hydrate and finally extracted with methylene chloride. The  
30 organic layer was dried over anhydrous sodium sulfate and evaporated to dryness to give 2.3 g of an oily product (93% yield).

35 Analogously the following products can be prepared starting from the suitable carboxylic acid:  
Methyl 4-ethoxy butanoate;  
Methyl cyclopropylacetate; and  
5-methoxycarbonylethyl-2-amino-1,3-thiazole.

**Example 11****4-methyl-pentanal**

1.24 ml (14.18 mmol) of oxalyl chloride were dissolved in  
5 10 ml of methylene chloride and after cooling to -60°C,  
2.31 ml of DMSO (35 mmoles) were dropped.  
After 5 minutes at the same temperature, a solution of 1 ml  
(11.9 mmol) of 4-methyl-1-pentanol in 10 ml of methylene  
10 chloride was slowly dropped. The mixture was maintained  
under stirring for 30 minutes at the same temperature, then  
8.3 ml (59.5 mmol) of triethylamine were added. After 2  
hours at 0°C water was added. The mixture was diluted with  
methylenec chloride and washed successively with 1M  
15 hydrochloric acid, water, saturated sodium bicarbonate and  
finally with brine. The organic layer was dried over  
anhydrous sodium sulfate and evaporated to dryness to give  
0.7 g (25% yield) of the title compound.  
Analogously the following products can be prepared starting  
from the suitable alcohol:  
20 2-cyclopropyl-1-ethanal;  
4-methylthio-1-butanal;  
4-ethoxy-1-butanal;  
5-methoxy-1-pentanal; and  
5-dimethylaminoimino-1-hexanal.

25

**Example 12****5-benzyloxy-1-methoxy-pentane**

1.6 g (0.039 mol) of 55% sodium hydride in oil were added to  
50 ml of dimethylformamide under stirring at room  
30 temperature. 5 ml (0.026 mol) of 5-benzyloxy-1-pentanol and  
2.43 ml (0.039 mol) of methyl iodide were then added  
successively. After a night the excess of sodium hydride  
was decomposed with water and the solvent evaporated under  
vacuum. The residue was redissolved with methylene  
35 chloride and washed with water. The organic layer was  
finally dried over anhydrous sodium sulfate and evaporated  
to give 3.5 g (70% yield) of the title compound.

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Analogously, by using ethyl iodide, the following compound can be prepared:

4-ethoxy-butanoic acid.

5 **Example 13**

**5-methoxy-1-pentanol**

3.5 g (0.018 mol) of 5-benzyloxy-1-methoxy-pentane were dissolved in 50 ml of ethanol and 400 mg of 10% palladium on activated charcoal were added. The mixture was hydrogenated at 40 psi at room temperature for 5 hours, then filtered on celite and evaporated under vacuum to give 1.77 g (84% yield) of the title compound.

**Example 14**

**Ethyl 5-dimethylaminoimino-hexanoate**

15.8 g (100 mmol) of ethyl 4-acetyl-butanoate and 6 g (100 mmol) of anhydrous N,N-dimethyl hydrazine in 50 ml of toluene containing 0.1 ml of trifluoroacetic acid were heated at 70 °C for 5 hours. The mixture was then washed with water, dried over anhydrous sodium sulfate and evaporated to give 12.3 g (79% yield) of the title compound.

**Example 15**

**N-[5-(3-oxo-butyl)-1,3-thiazol-2-yl]-butanamide**

25 To a stirred solution of 200 mg (1 mmol) of cupric acetate in 10 ml of water 141 mg (0.5 mmol) of N-[5-(3-dimethylaminoimino-butyl)-1,3-thiazol-2-yl]-butanamide in 10 ml of tetrahydrofuran were added. After 2 hours the solvent was removed under reduced pressure, a mixture of aqueous ammonium chloride and ammonium hydroxide was added and the product extracted with methylene chloride to give after drying and concentration 114 mg (95% yield) of the title compound.

35

**Example 16**

**2-benzyloxycarbonylamino-5-formyl-1,3-thiazole**

1 g (7.8 mmol) of 2-amino-5-formyl-1,3-thiazole was dissolved in 25 ml of tetrahydrofuran and 1.35 ml (9.36 mmol) of triethylamine and 1.33 ml (9.36 mmol) of benzylchloroformate were added at 0°C under stirring. After 5 8 hours at room temperature the solvent was evaporated, the residue redissolved with methylene chloride and washed with saturated tartaric acid and then with water. The solvent was dried over anhydrous sodium sulfate and evaporated. The residue was purified by chromatography on silica gel using 10 cyclohexane-ethylacetate as eluent to give 1.3 g (65% yield) of the title compound.

**Example 17****5-hydroxymethyl-2-benzylloxycarbonylamino-1,3-thiazole**

15 530 mg (14 mmol) of sodium borohydride were added in small portions to a stirred solution of 7 g (27 mmol) of 2-benzylloxycarbonylamino-5-formyl-1,3-thiazole in 80 ml of methanol at room temperature. The reaction went on for 2 hours. After evaporation of the solvent the residue was 20 purified by chromatography (cyclohexane-ethylacetate) to give 5.05 g (71% yield) of the title compound.

**Example 18****2-benzylloxycarbonylamino-5-(4-phenyl-1-sulfonyloxy)methyl-1,3-thiazole**

25 To a solution of 1 g (3.78 mmol) of 2-benzylloxycarbonylamino-5-hydroxymethyl-1,3-thiazole in 25 ml of pyridine 0.86 g (4.54 mmol) of tosyl chloride in 10 ml of pyridine were dropped at 0°C. After stirring at room 30 temperature for 6 hours the solvent was evaporated under vacuum, the residue redissolved with methylene chloride, washed with 1M hydrochloric acid and finally with water. The organic layer was dried over anhydrous sodium sulfate and evaporated. The residue was purified by chromatography 35 on silica gel (cyclohexane-ethylacetate) to give 1.2 g (80% yield) of the title compound.

**Example 19**

**2-benzyloxycarbonylamino-5-(2-ethoxycarbonyl-3-ethoxycarbonylethyl)-1,3-thiazole**

To a suspension of 321 mg of 55% sodium hydride in oil (7.4 mmol) in 20 ml of tetrahydrofuran 1.12 ml (7.4 mmol) of

5 diethylmalonate were added. After 30 minutes, a solution of  
 1.5 g (3.7 mmol) of 2-benzyloxycarbonylamino-5-(4-phenyl-1-  
 sulphonyloxy)methyl-1,3-thiazole in 10 ml of the same  
 solvent was dropped under stirring. After 6 hours the  
 solvent was evaporated and the residue redissolved with  
 10 methylene chloride and washed with water. The organic layer  
 was dried over anhydrous sodium sulfate and evaporated. The  
 residue was chromatographed on a silica gel column  
 (cyclohexane-ethylacetate) to give 1.05 g (70% yield) of  
 the title compound.

15

**Example 20**

$$2\text{-benzylxycarbonylamino-5-ethoxycarbonyl} \text{ethyl-1,3-thiazole}$$

To a solution of 4.06 g (10 mmol) of 2-benzylloxycarbonylamino-5-(2-ethoxycarbonyl-3-

20 ethoxycarbonylethyl)-1,3-thiazole in 10 ml of dimethylsulphoxide 0.64 g (11 mmol) of sodium chloride and 0.36 (20 mmol) of water were added under stirring. The mixture was heated at 160 °C for 8 hours and then the solvent removed under vacuum. The residue was redissolved 25 with methylene chloride and washed with brine. After drying and concentration the residue was chromatographed on a silica gel column (cyclohexane-ethylacetate) to give 2.67 g (80% yield) of the title compound.

### 30 Example 21

### 2-amino-5-carboxyethyl-1,3-thiazole

1 g (2.9 mmol) of 2-benzyloxycarbonylamino-5-ethoxycarbonylethyl-1,3-thiazole was dissolved in 20 ml of 33% hydrobromic acid in acetic acid. After 2 hours at room temperature, the solvent was evaporated under vacuum. The residue was redissolved in the minimum amount of water and the hydrobromide of the title compound was precipitated by adding diethylether (75% yield).

**Example 22****Preparation of methyl 2-[3-(3-chloropropoxy)phenyl]acetate**

5 A mixture of methyl (m-hydroxyphenyl)acetate ((5 g, 0.03 moles), 1-bromo-3-chloropropane (3.26 ml, 0.03 moles) and anhydrous potassium carbonate (6.4 g) in anhydrous acetone (60 ml) was refluxed for 40 hours. After cooling, the precipitate was filtered off and the solution was evaporated to dryness to give the product as an oil, which  
10 was purified by flash chromatography with hexane:AcOEt (97:3) as eluent (6.2 g, 85% yield).

Analogously, the following product can be prepared:  
methyl 2-[3-(2-chloroethoxy)phenyl]acetate.

**15 Example 23****Preparation of 2-[3-(3-chloropropoxy)phenyl]acetic acid**

20 A mixture of methyl 2-[3-(3-chloropropoxy)phenyl]acetate (4.95 g, 0.02 moles) and a solution of 1N sodium hydroxide (0.02 moles) was stirred at room temperature for 24 hours. After acidification the acid separated as white powder (4.53 g, 97% yield)  
m.p. 83-84°C  
Analogously, the following product can be prepared:  
2-[3-(2-chloroethoxy)phenyl]acetic acid  
25 m.p. 100-101°C.

**Example 24****Preparation of N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-[3-(4-morpholinyl)propoxy]phenyl]acetamide**

30 A mixture of 2-[3-(3-chloropropoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide (1.00 g, 2.8 mmoles), morpholine (1.24 ml, 14.2 mmoles), potassium iodide (0.24 g, 1.4 mmoles) in anhydrous dimethylformamide (3.5 ml) was heated at 100°C for 6 hours. The solution was acidified and extracted with ether to eliminate unreacted products; then the solution was basified and extracted with ether. The solvent was evaporated to dryness to give the product as an oily semisolid which was purified by flash chromatography

with dichloromethane:methanol (97:3) as eluent (1.0 g, 87% yield)

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.09 (bs, 1H, CONH); 7.21 (t, 1H, H5 Ph); 7.13 (s, 1H, thiazole CH); 6.8-6.9 (m, 3H, H2, H4, H6 Ph); 3.97 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N); 3.66 (s, 2H, COCH<sub>2</sub>); 3.54 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>N); 3.07 (ept, 1H, CHMe<sub>2</sub>); 2.39 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N); 2.33 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>N); 2.14 (quint, 2H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N); 1.22 (d, 6H, CHMe<sub>2</sub>).

10 Analogously, the following product can be prepared:

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-[2-(4-morpholinyl)ethoxy]phenyl)acetamide

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.11 (bs, 1H, CONH); 7.20 (t, 1H, H5 Ph); 7.13 (d, 1H, thiazole CH); 6.7-6.9 (m, 3H, H2, H4, H6 Ph); 4.04 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>N); 3.66 (s, 2H, COCH<sub>2</sub>); 3.55 (m, 4H, OCH<sub>2</sub>CH<sub>2</sub>N morpholine); 3.08 (m, 1H, CHMe<sub>2</sub>); 2.66 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>N); 2.44 (m, 4H, OCH<sub>2</sub>CH<sub>2</sub>N morpholine); 1.22 (d, 6H, CHMe<sub>2</sub>);

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-[3-(1-pyrrolidinyl)propoxy]phenyl)acetamide

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.1 (bs, 1H, CONH); 7.19 (t, 1H, H5 Ph); 7.13 (d, 1H, thiazole CH); 6.7-6.9 (m, 3H, H2, H4, H6 Ph); 3.97 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N); 3.66 (s, 2H, COCH<sub>2</sub>); 3.08 (m, 1H, CHMe<sub>2</sub>); 2.50 (m, 2H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N); 2.41 (m, 4H, CH<sub>2</sub>N pirrolidine); 1.85 (m, 2H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N); 1.65 (m, 4H, CH<sub>2</sub>CH<sub>2</sub>N pirrolidine); 1.23 (d, 6H, CHMe<sub>2</sub>);

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-[3-(4-methyl-1-piperazinyl)propoxy]phenyl)acetamide

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.1 (bs, 1H, CONH); 7.19 (t, 1H, H5 Ph); 7.13 (d, 1H, thiazole CH); 6.7-6.9 (m, 3H, H2, H4, H6 Ph); 3.95 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N); 3.66 (s, 2H, COCH<sub>2</sub>); 3.08 (m, 1H, CHMe<sub>2</sub>); 2.15-2.45 (m, 10H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N+piperazine); 2.11 (s, 3H, NMe); 1.82 (m, 2H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N); 1.22 (d, 6H, CHMe<sub>2</sub>); 2-(3-[2-(dimethylamino)ethoxy]phenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.08 (bs, 1H, CONH); 7.2-6.90 (m,

5H, Ph+thiazole CH); 4.00 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>N); 3.66 (s, 2H, COCH<sub>2</sub>); 3.07 (m, 1H, CHMe<sub>2</sub>); 2.59 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>N); 2.11 (s, 3H, NMe); 2.19 (s, 6H, Me<sub>2</sub>N); 1.22 (d, 6H, CHMe<sub>2</sub>);  
 5 2-(3-[3-(dimethylamino)propoxy]phenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
 10 <sup>1</sup>H-NMR (DMSO-d<sub>6</sub>) δ ppm: 12.05 (bs, 1H, CONH); 7.19-6.79 (m, 5H, Ph+thiazole CH); 3.95 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N); 3.66 (s, 2H, COCH<sub>2</sub>); 3.08 (m, 1H, CHMe<sub>2</sub>); 2.32 (t, 2H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N); 2.11 (s, 3H, NMe<sub>2</sub>); 1.81 (m, 2H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N); 1.22 (d, 6H, CHMe<sub>2</sub>).

**Example 25**

**Preparation of 2-[N-[2'-N'-(ethoxycarbonyl-methyl)-amino]-acetyl]-amino-5-bromo-thiazole**

15 A solution of N-(5-bromo-1,3-thiazol-2-yl)-2-bromoacetamide (0.35 g, 1.17 mmol) in DMF (5 ml) was added dropwise to a solution of glycine ethyl ester hydrochloride (0.33 g, 2.33 mmol) and triethylamine (0.49 ml, 3.5 mmol) in DMF (10 ml). After 3 hours at room temperature, the reaction mixture was  
 20 heated at 40°C for about 5 hours and then diluted with water and extracted with methylene chloride. The combined organic layers were washed with brine, dried, concentrated and chromatographed on silica gel using cyclohexane:ethyl acetate 7:3 as eluent. The title compound was obtained as a  
 25 colourless solid (0.15 g, 43%)  
 m.p. 115-116°C

<sup>1</sup>H-NMR (DMSO-d<sup>6</sup>) δ ppm: 7.46 (s, 1H, H<sub>4</sub>thiaz), 4.05 (q, 2H, OCH<sub>2</sub>CH<sub>3</sub>), 3.49 (s, 2H, NHCOCH<sub>2</sub>), 3.4 (s, 2H, NHCH<sub>2</sub>), 1.18 (t, 3H, OCH<sub>2</sub>CH<sub>3</sub>).  
 30

Analogously, the following compound can be prepared:

2-anilino-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide: m.p. 143-145°C

<sup>1</sup>H-NMR (DMSO-d<sup>6</sup>) δ ppm: 11.92 (s, 1H, NHCO), 7.13 (s, 1H, H<sub>4</sub>thiaz), 7.06-6.6 (m, 5H, Ph), 6.0 (t, 1H, NHCH<sub>2</sub>), 3.95 (d, 2H, NHCH<sub>2</sub>), 3.08 (m, 1H, CHMe<sub>2</sub>), 1.23 (d, 6H, CHMe<sub>2</sub>).

**Example 26****Preparation of N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-bromophenyl)acetamide**

5 To a suspension of resin N-Cyclohexylcarbodiimide N'-methylpolystyrene (0.251 g, 2.39 mmol g<sup>-1</sup>, 0.6 mmol), previously washed with DCM (3X5 ml), in DCM (4 ml) at room temperature, 2-bromophenylacetic acid (0.086 g, 0.4 mmol) was added. After 10 min., a solution of 2-amino-5-isopropyl-1,3-thiazole (0.0284 g, 0.2 mmol) in DCM (4 ml) was added. The mixture was shaked for 24 hours at room temperature, the resin filtered and washed with DCM (3X10 ml). The filtrated were combined, washed with water, 5% HCl, water, saturated sodium bicarbonate and water, dried over sodium sulfate and evaporated.

10 <sup>1</sup>H-NMR (DMSO-d<sup>6</sup>) δ ppm: 10.05 (s broad, 1H, NHCOCH<sub>2</sub>), 7.6-7.2 (m, 4H, Ar), 7.08 (s, 1H, H4thiaz), 3.98 (s, 2H, NHCOCH<sub>2</sub>), 3.11 (m, 1H, CHMe<sub>2</sub>), 1.31 (d, 6H, CHMe<sub>2</sub>)

15 20 All the compounds were characterised by Mass Spectroscopy (MS). LC-MS confirmed that in each case the principle component had a molecular ion corresponding to the expected product.

25 Chromatography: Reverse phase HPLC with UV detection were run.  
Mobile A: water (0.1% TFA)  
Mobile B: acetonitrile:water 95:5 (0.1% TFA)  
Flow rate: 1ml/min

30 Gradient: 10-100% B in 12 minutes, hold 100% B 3 min, return 10% B in 5 min  
Detection: UV monitor 215, 254 and 300 nm  
Sample were prepared as dilute solutions in acetonitrile (1-1.5 mM).

35 The compounds showed an HPLC area % ranging from 40 to 100%.

Starting from the suitable carboxylic acid, the following compounds can be prepared:

5 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,3,4,5,6-  
pentafluorophenyl)acetamide;

10 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-chlorophenyl)acetamide;

15 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-nitrophenyl)acetamide;

20 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-trifluoromethylphenyl)acetamide;

25 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-methoxyphenyl)acetamide;

30 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-dimethoxyphenyl)acetamide;

35 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-difluorophenyl)acetamide;

40 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4,5-trimethoxyphenyl)acetamide;

45 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,6-dichlorophenyl)acetamide;

50 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-chloro-6-fluorophenyl)acetamide;

55 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,5-dimethoxyphenyl)acetamide;

60 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,5-difluorophenyl)acetamide;

65 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-bis-trifluoromethylphenyl)acetamide;

70 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methylthiophenyl)acetamide;

75 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide;

80 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-bromophenyl)acetamide;

85 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-chlorophenyl)acetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl)acetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-trifluoromethylphenyl)acetamide;

5 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methylphenyl)acetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-trifluoromethylphenyl)acetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-chlorophenyl)acetamide;

10 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-dinitrophenyl)acetamide;

15 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-dichlorophenyl)acetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-difluorophenyl)acetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-benzyloxy-3-methoxyphenyl)acetamide;

20 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dichlorophenyl)acetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-difluorophenyl)acetamide;

25 2-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide;

2-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2,2-diphenylacetamide;

30 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-nitrophenoxy)acetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl)propanamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-isobutylphenyl)propanamide;

35 N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxo-2-phenylacetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-phenylpentanamide;

(E, Z)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-2-butenamide;

N-(5-isopropyl-1,3-thiazol-2-yl)bicyclo[4.2.0]octa-1,3,5-triene-7-carboxamide;

5 N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylbutanamide; tert-butyl (1R)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethylcarbamate; (1R)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate;

10 (1S)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate; 2-(acetylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide;

(R)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide;

15 3,3,3-trifluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxy-2-phenylpropanamide; 2-(2,4-dinitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;

20 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(5-benzyloxy-1H-indol-3-yl)acetamide; N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxy-2-methyl-1H-indol-3-yl)acetamide; 2-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-

25 oxoacetamide; 2-[1-(4-chlorobenzoyl)-5-methoxy-2-methyl-1H-indol-3-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide; 4-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)butanamide;

30 N-(5-isopropyl-1,3-thiazol-2-yl)-3-(2-thienyl)propanamide 2-(5-chloro-1-benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide; 2-(1-benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;

35 2-[2-(formylamino)-1,3-thiazol-4-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamide; 2-(2-[(2-chloroacetyl)amino]-1,3-thiazol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamide;

2-chloro-N-(4-(2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl)-1,3-thiazol-2-yl)acetamide;  
ethyl 2-({[2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-(1H-pyrazol-3-yl)ethylidene]amino}oxy)acetate;

5 N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxo-4-(4-methyl-phenyl)butanamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-nitrophenyl)butanamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenylbutanamide;

10 benzyl 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]-4-oxobutylcarbamate;  
4-(1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)butanamide;

15 N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-methoxy-1-naphthyl)-4-oxobutanamide;  
3-(2-chlorophenoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide;  
3-(4-methylphenoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide;

20 3-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide;  
3-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylpentanamide;  
3-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide;

25 3-(4-methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide;  
3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide;  
3-phenyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-5-oxo-5-phenylpentanamide;

30 2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate;  
N-(5-isopropyl-1,3-thiazol-2-yl)-2-[4-(1-oxo-1,3-dihydro-2H-isoindol-2-yl)phenyl]propanamide;

35 1-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)cyclopentanecarboxamide;  
1-phenyl-N-(5-isopropyl-1,3-thiazol-2-yl)cyclopentanecarboxamide;

2-(3-bromo-4-methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;

2-(2-nitro-4-trifluoromethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;

5 5-cyclohexyl 1-(4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}benzyl) (2S)-2-[(tert-butoxycarbonyl)amino]pentanedioate;

2-(5,6-dimethyl-1H-benzimidazol-1-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;

10 2-[5-(4-chlorophenyl)-2H-1,2,3,4-tetraazol-2-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-[5-(1-pyrrolidinyl)-2H-1,2,3,4-tetraazol-2-yl]acetamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methyl-1-benzothiophen-2-yl)acetamide;

15 N-(5-isopropyl-1,3-thiazol-2-yl)-4,4-bis(4-methylphenyl)-3-butenamide;

2-cyclopropyl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;

N-(4-bromo-6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexyl)benzamide;

20 2-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;

benzyl 6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexylcarbamate;

N-1-(5-isopropyl-1,3-thiazol-2-yl)-N-4-(2-propynyl)-2-butenediamide;

25 4-(2,4-dimethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide;

4-(4-benzyloxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide;

30 4-(thiphen-2-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide;

benzyl 2-[(benzyloxy)carbonyl]amino]-5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate;

4-(1H-indol-3-yl)-N-(3-[(5-isopropyl-1,3-thiazol-2-yl)amino]-3-oxopropyl)butanamide;

35 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonylphenyl 4-chlorobenzenesulfonate;

N- (5-isopropyl-1,3-thiazol-2-yl)-4-{[(2-methoxyanilino)carbonyl]amino}benzamide;  
4-[(2-(isopropylsulfonyl)acetyl]amino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

5 N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(2-phenylsulfanyl)acetyl]amino}benzamide;  
4-[(diethylamino)sulfonyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
2-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

10 3,5-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
3-[(2-fluoroanilino)carbonyl]amino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-1-phenyl-5-propyl-1H-pyrazole-4-carboxamide;

15 3-chloro-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfanyl)-2-thiophenecarboxamide;  
3-iodo-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfanyl)-2-thiophenecarboxamide;  
2-[(4-chlorophenyl)sulfonyl]methyl)-N-(5-isopropyl-1,3-

20 thiazol-2-yl)-4-methyl-1,3-thiazole-5-carboxamide;  
5-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(trifluoromethyl)-3-furamide;  
3,5-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
3,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

25 2,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
2,3-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
3-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
2-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

30 4-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
3-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
4-chloro-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

35 5-bromo-2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

2,4-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

3,4-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
2,3,4,5,6-pentafluoro-N-(5-isopropyl-1,3-thiazol-2-  
y1)benzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-3-nitrobenzamide;  
5 N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-nitrobenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-nitrobenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-  
nitrobenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxy-2-  
10 nitrobenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-2-  
nitrobenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxy-3-  
nitrobenzamide;  
15 N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-  
nitrobenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dinitrobenzamide;  
5-{{(5-isopropyl-1,3-thiazol-2-yl)amino}carbonyl}-2-  
nitrophenyl octanoate;  
20 N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamide;  
4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide;  
4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide;  
2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamide;  
25 5-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide;  
2-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-5-nitrobenzamide;  
4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide;  
4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitro-4-  
30 (trifluoromethyl)benzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-  
bis(trifluoromethyl)benzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-2,6-  
bis(trifluoromethyl)benzamide;  
35 N-(5-isopropyl-1,3-thiazol-2-yl)-2-  
(trifluoromethyl)benzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-3-  
(trifluoromethyl)benzamide;

3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethyl)benzamide;

2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide;

5 5-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide;

2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethyl)benzamide;

4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide;

10 methyl 2-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)benzoate;

4-cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

3-cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

15 N-(5-isopropyl-1,3-thiazol-2-yl)-3-methylbenzamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-methylbenzamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylbenzamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-4-vinylbenzamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-4-(2-

20 phenylethynyl)benzamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-methylbenzamide;

2-benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenethylbenzamide;

25 N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylbenzamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenylbenzamide;

4-(tert-butyl)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-4-isopropylbenzamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-4-pentylbenzamide;

30 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylbenzamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-3,4-dimethylbenzamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethylbenzamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-4-

35 (methylsulfonyl)benzamide;

3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide;

3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide;  
5-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxybenzamide;

5 N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxybenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxybenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-3,4-dimethoxybenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethoxybenzamide;

10 N-(5-isopropyl-1,3-thiazol-2-yl)-2,4-dimethoxybenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-2,3-dimethoxybenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-3-phenoxybenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenoxybenzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenoxybenzamide;

15 2-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
4-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-3,4,5-trimethoxybenzamide;  
3,4-diethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
3,4,5-triethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

20 N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-(methoxymethoxy)benzamide;  
4-butoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-4-propoxybenzamide;  
4-isopropoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

25 N-(5-isopropyl-1,3-thiazol-2-yl)-1,3-benzodioxole-5-carboxamide;  
4-(benzyloxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
4-(2-cyclohexen-1-yloxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

30 N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethoxy)benzamide;  
4-(difluoromethoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfanyl)benzamide;

35 2-[(4-chlorophenyl)sulfinyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

N- (5-isopropyl-1,3-thiazol-2-yl)-2-[(4-nitrophenyl)sulfinyl]benzamide;

N- (5-isopropyl-1,3-thiazol-2-yl)-4-[(4-methylphenyl)sulfonyl]-3-nitrobenzamide;

5 N- (5-isopropyl-1,3-thiazol-2-yl)-3-[(trifluoromethyl)sulfanyl]benzamide;

N- (5-isopropyl-1,3-thiazol-2-yl)-2-methoxy-4-(methylsulfanyl)benzamide;

10 2-[(2-cyanophenyl)sulfanyl]-N- (5-isopropyl-1,3-thiazol-2-yl)benzamide;

N-1~,N-1~-diethyl-3,6-difluoro-N~2~- (5-isopropyl-1,3-thiazol-2-yl)phthalamide;

4-formyl-N- (5-isopropyl-1,3-thiazol-2-yl)benzamide;

2-formyl-N- (5-isopropyl-1,3-thiazol-2-yl)benzamide;

15 4-{[(2,5-dimethoxyanilino)carbonyl]amino}-N- (5-isopropyl-1,3-thiazol-2-yl)benzamide;

4-(hydroxymethyl)-N- (5-isopropyl-1,3-thiazol-2-yl)benzamide;

4-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-2-

20 nitrobenzyl acetate;

4-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-2-nitrobenzyl 4-(acetylamino)-3-iodobenzoate;

4-(acetylamino)-N- (5-isopropyl-1,3-thiazol-2-yl)benzamide;

N- (5-isopropyl-1,3-thiazol-2-yl)-4-[(2-

25 phenylacetyl)amino]benzamide;

4-(acetylamino)-3-iodo-N- (5-isopropyl-1,3-thiazol-2-yl)benzamide;

4-amino-N- (5-isopropyl-1,3-thiazol-2-yl)benzamide;

4-(dimethylamino)-N- (5-isopropyl-1,3-thiazol-2-

30 yl)benzamide;

3-(dimethylamino)-N- (5-isopropyl-1,3-thiazol-2-yl)benzamide;

2-(methylamino)-N- (5-isopropyl-1,3-thiazol-2-yl)benzamide;

N- (5-isopropyl-1,3-thiazol-2-yl)-2-[3-

35 (trifluoromethyl)anilino]benzamide;

3-{[(5-bromo-1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)methyl]amino}-N- (5-isopropyl-1,3-thiazol-2-yl)benzamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-4-(1H-pyrrol-1-yl)benzamide;

2,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)isonicotinamide;

5 2-(4-bromophenyl)-6-(4-iodophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)isonicotinamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(trifluoromethyl)anilino]nicotinamide;

5,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide;

10 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-6-methylnicotinamide;

2,6-dichloro-5-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenoxy nicotinamide;

15 N-(5-isopropyl-1,3-thiazol-2-yl)-6-(2,2,2-trifluoroethoxy)nicotinamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2,6-dimethoxy nicotinamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-quinoxalinecarboxamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-

20 pyrazinecarboxamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-8-quinolinecarboxamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-4-quinolinecarboxamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-1-phenyl-1H-

25 pyrazole-4-carboxamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-1H-pyrazole-3-carboxamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-1H-pyrazole-4-carboxamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-phenyl-2H-

30 1,2,3-triazole-4-carboxamide;

2-[(2,1,3-benzoxadiazol-5-yloxy)methyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-1,3-thiazole-5-carboxamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-9H-fluorene-1-carboxamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-7-methoxy-1-benzofuran-2-

35 carboxamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-1-[(4-methylphenyl)sulfonyl]-1H-pyrrole-3-carboxamide;

2-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamide;

4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-2-naphthamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-9,10-dioxo-9,10-dihydro-2-anthracenecarboxamide;

5 N-(5-isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluorene-4-carboxamide  
N-(5-isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluorene-1-carboxamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-8-oxo-5,6,7,8-tetrahydro-10-2-naphthalenecarboxamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-1,3-dioxo-1,3-dihydro-2-benzofuran-5-carboxamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-5-carboxamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-4-carboxamide;

15 N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-2-phenyl-1H-indole-5-carboxamide;  
2-butyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indole-5-carboxamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-6-carboxamide;

20 N-(5-isopropyl-1,3-thiazol-2-yl)-5-methoxy-1H-indole-2-carboxamide;  
1-allyl-2-butyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-5-carboxamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indole-2-carboxamide;

25 1-benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-1H-indole-5-carboxamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-1H-1,2,3-benzotriazole-5-carboxamide;

30 N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-isoxazolecarboxamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-3-thiophenecarboxamide;  
N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-thiophenecarboxamide;

35 N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-thiophenecarboxamide;  
5-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-2-thiophenecarboxamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-3-[(2,3,3-trichloroacryloyl)amino]-2-thiophenecarboxamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-5-(4-nitrophenyl)-2-furamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-5-(2-nitrophenyl)-2-furamide;

5-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-5-[3-(trifluoromethyl)phenyl]-2-furamide;

5-(4-chloro-2-nitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide;

N-(5-isopropyl-1,3-thiazol-2-yl)-5-(4-methyl-2-nitrophenyl)-2-furamide

5-[2-chloro-5-(trifluoromethyl)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide;

tert-butyl (1S)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-1-methyl-2-oxoethylcarbamate

tert-butyl (1S,2S)-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]-2-methylbutylcarbamate

tert-butyl 2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

tert-butyl (1S)-5-amino-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonylpentylcarbamate

tert-butyl 4-[(imino{[(4-methylphenyl)sulfonyl]amino}methyl)amino]-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonylbutylcarbamate

tert-butyl 1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl-3-(tritylamo)propylcarbamate

tert-butyl (1S)-1-(benzyloxymethyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

tert-butyl (1S)-1-benzyl-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

tert-butyl (1R)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-(benzylthiomethyl)ethylcarbamate

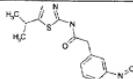
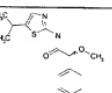
benzyl (3S)-3-[(tert-butoxycarbonyl)amino]-4-[(5-isopropyl-1,3-thiazol-2-yl)amino]-4-oxobutanoate

tert-butyl (2S)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]-1-pyrrolidinecarboxylate  
 tert-butyl (1S)-1-(1H-indol-3-ylmethyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate  
 5 tert-butyl (1S)-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)-3-(methylsulfanyl)propylcarbamate  
 tert-butyl (1S)-2-benzyloxy-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonylpropylcarbamate  
 tert-butyl (1S)-1-(4-benzyloxybenzyl)-2-[(5-isopropyl-1,3-  
 10 thiazol-2-yl)amino]-2-oxoethylcarbamate  
 tert-butyl (1S)-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)-2-methylpropylcarbamate  
 tert-butyl (1S)-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)-3-methylbutylcarbamate and  
 15 benzyl (4S)-4-[(tert-butoxycarbonyl)amino]-5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate.

Following the same procedure as reported in Example 3, the compounds described in the table (I) below can be prepared:

20

Table I

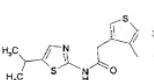
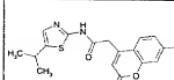
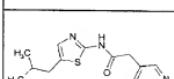
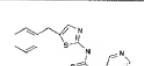
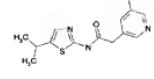
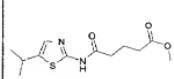
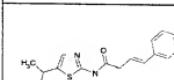
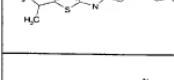
MOLSTRUCTURE	m p °C	<sup>1</sup> H-NMR	Sol vent
		12.23 (s broad, 1H, <u>NHCOCH<sub>2</sub></u> ), 8.22-7.62 (m, 4H, Ar), 7.15 (s, 1H, H4thiaz), 3.91 (s, 2H, NHCOCH <sub>2</sub> ), 3.08 (m, 1H, <u>CHMe<sub>2</sub></u> ), 1.22 (d, 6H, <u>CHMe<sub>2</sub></u> )	DMSO-d <sup>6</sup>
		9.81 (s broad, 1H, <u>NHCOCH<sub>2</sub></u> ), 7.5- 7.3 (m, 4H, Ar), 7.11 (s, 1H, H4thiaz), 4.83 (s, 1H, NHCOCH), 3.44 (s, 3H, Ome) 3.11 (m, 1H, CHMe <sub>2</sub> ), 1.3 (d, 6H, <u>CHMe<sub>2</sub></u> )	DMSO-d <sup>6</sup>

	124 - 125	12.06 (s broad, 1H, NHCO), 7.13 (s, 1H, H4thiaz) 6.92-6.81 (m, 3H, Ar), 3.72 (s, 3H, OMe), 3.70 (s, 3H, OMe), 3.61 (s, 2H, NHCOCH <sub>2</sub> ), 3.07 (m, 1H, CHMe <sub>2</sub> ), 1.22 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	77- 78	12.05 (s broad, 1H, NHCO), 7.38-7.29 (m, 5H, Ar), 7.12 (s, 1H, H4thiaz), 4.95 (s, 1H, CHOMe), 3.23 (s, 2H, CHOMe), 3.05 (m, 1H, CHMe <sub>2</sub> ), 1.20 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	136 - 137	12.08 (s broad, 1H, NHCOCH <sub>2</sub> ), 7.28 (d, 2H, Ar), 7.13 (s, 1H, H4thiaz), 7.1 (d, 2H, Ar), 3.65 (s, 2H, NHCOCH <sub>2</sub> ), 3.06 (m, 1H, CHMe <sub>2</sub> ), 2.98 (s, 6H, NM <sub>2</sub> ), 1.22 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	130 - 131	12.22 (s, 1H, NHCO), 7.85-7.48 (m, 7H, Ar), 7.14 (s, 1H, H4thiaz), 3.89 (s, 2H, CH <sub>2</sub> CO), 3.07 (m, 1H, CHMe <sub>2</sub> ), 1.22 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	130 - 131	12.16 (s, 1H, NHCO), 7.52-7.29 (m, 4H, Ar), 7.14 (s, 1H, H4thiaz), 3.73 (s, 2H, CH <sub>2</sub> CO), 3.08 (m, 1H, CHMe <sub>2</sub> ), 1.22 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	177 - 178	8.07-7.48 (m, 7H, Ar), 7.15 (s, 1H, H4thiaz), 4.22 (s, 2H, CH <sub>2</sub> CO), 3.06 (m, 1H, CHMe <sub>2</sub> ), 1.20 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	223 - 224	12.61 (s, 1H, NHCO), 7.69-7.51 (m, 4H, Ar), 7.19 (s, 1H, H4thiaz), 4.55 (dd, 1H, CHCO), 3.08 (m, 1H, CHMe <sub>2</sub> ), 2.89 (m, 2H, COCH <sub>2</sub> CH), 1.22 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	105 - 106	12.50 (s, 1H, NHCO), 7.53-7.51 (m, 5H, Ar), 7.18 (s, 1H, H4thiaz), 6.12 (d, 1H, J <sub>H-F</sub> = 46.8, CHF), 3.09 (m, 1H, CHMe <sub>2</sub> ), 1.22 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	150 - 152	11.20 (s broad, 1H, NHCO), 7.28-7.07 (m, 5H, Ar+H4thiaz), 3.80 (s, 2H, CH <sub>2</sub> CO), 3.13 (m, 1H, CHMe <sub>2</sub> ), 1.32 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>

<chem>CN(C)c1nc2sc(C(=O)c3ccc(F)cc3)nc2s1</chem>	164 - 166	11.45 (s broad, 1H, NHCO), 7.37-7.14 (m, 5H, Ar+H4thiaz), 3.88 (s, 2H, NHCOCH <sub>2</sub> ), 3.12 (m, 1H, CHMe <sub>2</sub> ), 1.32 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
<chem>CN(C)c1nc2sc(C(=O)c3ccc(F)cc3)nc2s1</chem>	98-100	8.35 (s broad, 1H, NHCO), 7.40 (m, 5H, Ar), 6.99 (s, 1H, H4thiaz), 3.10 (m, 1H, CHMe <sub>2</sub> ), 1.78 (m, 2H, CH <sub>2</sub> ), 1.29 (m, 2H, CH <sub>2</sub> ), 1.25 (d, 6H, CHMe <sub>2</sub> )	CDCl <sub>3</sub>
<chem>CC=C[C@H]1[C@H](COC(=O)C)C[C@H](C1)C(=O)N2C[C@H]3[C@H]2C[C@H]3C=C(C)C</chem>	130 - 132	12.06 (s broad, 1H, NHCOCH <sub>2</sub> ), 7.13 (s, 1H, H4thiaz), 6.86-6.75 (m, 3H, Ar), 5.96 (s, 2H, OCH <sub>2</sub> O), 3.60 (s, 2H, NHCOCH <sub>2</sub> ), 3.05 (m, 1H, CHMe <sub>2</sub> ), 1.22 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
<chem>CN(C)c1nc2sc(C(=O)c3ccc(F)cc3)nc2s1</chem>	100 - 102	12.1 (s broad, 1H, NHCOCH <sub>2</sub> ), 7.2-7 (m, 4H, Ar+H4thiaz), 3.64 (s, 2H, NHCOCH <sub>2</sub> ), 3.07 (m, 1H, CHMe <sub>2</sub> ), 2.8-1.97 (m, 6H, -CH <sub>2</sub> CH <sub>2</sub> CH <sub>2</sub> -), 1.22 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
<chem>CN(C)c1nc2sc(C(=O)c3ccc(F)cc3)nc2s1</chem>	98-100	12.06 (s broad, 1H, NHCO), 7.3 (m, 5H, Ar), 7.03 (s, 1H, H4thiaz), 3.79 (q, 1H, CHMe), 3.10 (m, 1H, CHMe <sub>2</sub> ), 1.59 (d, 3H, CHMe), 1.30 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
<chem>CC=C[C@H]1[C@H](COC(=O)Cc2ccc(cc2)Nc3ccc(cc3)C)C[C@H](C1)C(=O)N2C[C@H]3[C@H]2C[C@H]3C=C(C)C</chem>	167 - 169	10 (s broad, 1H, NHCOCH <sub>2</sub> ), 7.6-7.4 (m, 9H, Ar), 7.04 (s, 1H, H4thiaz), 3.84 (s, 2H, NHCOCH <sub>2</sub> ), 3.11 (m, 1H, CHMe <sub>2</sub> ), 1.31 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
<chem>CC=C[C@H]1[C@H](COC(=O)Cc2ccc(cc2)Nc3ccc(cc3)C)C[C@H](C1)C(=O)N2C[C@H]3[C@H]2C[C@H]3C=C(C)C</chem>	115 - 116	12.06 (s broad, 1H, NHCO), 7.26 (m, 5H, Ar), 6.99 (s, 1H, H4thiaz), 3.79 (q, 1H, CHMe), 3.10 (m, 1H, CHMe <sub>2</sub> ), 1.59 (d, 3H, CHMe), 1.30 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
<chem>CC=C[C@H]1[C@H](COC(=O)Cc2ccc(cc2)Nc3ccc(cc3)C)C[C@H](C1)C(=O)N2C[C@H]3[C@H]2C[C@H]3C=C(C)C</chem>	112 - 114	12.06 (s broad, 1H, NHCO), 7.33 (m, 5H, Ar), 7.11 (s, 1H, H4thiaz), 3.93 (q, 1H, CHMe), 3.07 (m, 1H, CHMe <sub>2</sub> ), 1.40 (d, 3H, CHMe), 1.22 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
<chem>CC=C[C@H]1[C@H](COC(=O)Cc2ccc(cc2)Nc3ccc(cc3)C)C[C@H](C1)C(=O)N2C[C@H]3[C@H]2C[C@H]3C=C(C)C</chem>	124 - 126	12.01 (s broad, 1H, NHCO), 7.11-6-65 (m, 5H, Ar+H4thiaz), 3.55 (s, 2H, NHCOCH <sub>2</sub> ), 2.83 (s, 6H, NMe <sub>2</sub> ), 2.56 (d, 2H, CH <sub>2</sub> iPr), 1.74 (m, 1H, CHMe <sub>2</sub> ), 0.87 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>

	139 - 141	9.90 (s broad, 1H, NHCO), 7.04 (s, 1H, H4thiaz), 6.78 (m, 3H, Ar), 5.96 (s, 2H, OCH <sub>2</sub> O), 3.72 (s, 2H, NHCOCH <sub>2</sub> ), 2.60 (d, 2H, CH <sub>2</sub> iPr), 1.85 (m, 1H, CHMe <sub>2</sub> ), 0.93 (d, 6H, CHMe <sub>2</sub> )	CDCl <sub>3</sub>
	175 - 177	12.0 (s broad, 1H, NHCO), 7.28 (m, 6H, CH <sub>2</sub> Ph+H4thiaz), 7.08-6.64 (m, 4H, Ar), 4.04 (s, 2H, CH <sub>2</sub> Ph), 3.53 (s, 2H, NHCOCH <sub>2</sub> ), 2.82 (s, 6H, NMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	88-90	12.08 (s broad, 1H, NHCO), 7.20-6.81 (m, 5H, Ar+H4thiaz), 4.01 (dd, 2H, OCH <sub>2</sub> CH <sub>2</sub> OMe), 3.68 (s, 2H, NHCOCH <sub>2</sub> ), 3.61 (dd, 2H, OCH <sub>2</sub> CH <sub>2</sub> OMe), 3.3 (s, 3H, OCH <sub>2</sub> CH <sub>2</sub> OMe), 3.05 (m, 1H, CHMe <sub>2</sub> ), 1.22 (s, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	230 - 231	12.81 (s broad, 1H, NHCO), 8.63-7.79 (m, 3H, Ar), 7.71 (s, 2H, NH <sub>2</sub> ), 7.24 (s, 1H, H4thiaz), 3.12 (m, 1H, CHMe <sub>2</sub> ), 1.27 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	181 - 182	12.47 (s broad, 1H, NHCO), 8.13-7.37 (m, 4H, Ar), 7.23 (s, 1H, H4thiaz), 3.13 (m, 1H, CHMe <sub>2</sub> ), 1.27 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	263 - 264	12.0 (s broad, 1H, NHCO), 8.89-7.82 (m, 4H, Ar), 7.27 (s, 1H, H4thiaz), 3.13 (m, 1H, CHMe <sub>2</sub> ), 1.28 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	204 - 206	12.6 (s broad, 1H, NHCO), 8.06-7.60 (m, 3H, Ar), 7.23 (s, 1H, H4thiaz), 3.12 (m, 1H, CHMe <sub>2</sub> ), 1.27 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	148 - 150	8.54-8.31 (m, 3H, Ar), 6.98 (s, 1H, H4thiaz), 3.43 (s, 3H, SO <sub>2</sub> Me), 3.14 (m, 1H, CHMe <sub>2</sub> ), 1.35 (d, 6H, CHMe <sub>2</sub> )	CDCl <sub>3</sub>

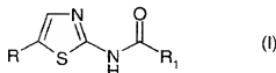
	173 - 175	8.16-8.06 (2d, 4H, Ar), 7.25 (s, 1H, H4thiaz), 3.88 (s, 3H, COOMe), 3.14 (m, 1H, CHMe <sub>2</sub> ), 1.28 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	164 - 166	8.50-7.86 (m, 3H, Ar), 7.24 (s, 1H, H4thiaz), 3.15 (m, 1H, CHMe <sub>2</sub> ), 1.28 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	178 - 179	12.4 (s broad, 1H, NHCO), 8.12-7.21 (m, 3H, Ar), 7.22 (s, 1H, H4thiaz), 3.2-2.48 (m, 5H, CHMe <sub>2</sub> , + piperazine), 2.22 (s, 3H, NMe), 1.27 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	170	12.6 (s broad, 1H, NHCO), 7.73-7.57 (m, 3H, Ar), 7.22 (s, 1H, H4thiaz), 3.15 (m, 1H, CHMe <sub>2</sub> ), 1.27 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	171	12.6 (s broad, 1H, NHCO), 8.16-8.05 (m, 4H, Ar), 7.24 (s, 1H, H4thiaz), 3.13 (m, 1H, CHMe <sub>2</sub> ), 2.62 (s, 3H, COMe), 1.28 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	207 - 209	9.4 (s broad, 1H, NHCO), 8.3 (s, 1H, NH), 7.55-6.98 (m, 6H, indole+H4thiaz), 3.96 (s, 2H, COCH <sub>2</sub> ), 3.10 (m, 1H, CHMe <sub>2</sub> ), 1.30 (d, 6H, CHMe <sub>2</sub> )	CDCl <sub>3</sub>
	116 - 118	9.80 (s broad, 1H, NHCO), 7.37-7.05 (m, 3H, Ar), 7.04 (d, 1H, H4thiaz), 3.84 (s, 2H, COCH <sub>2</sub> ), 3.11 (m, 1H, CHMe <sub>2</sub> ), 1.32 (d, 6H, CHMe <sub>2</sub> )	CDCl <sub>3</sub>
	148 - 150	10.20 (s broad, 1H, NHCO), 7.28-7.01 (m, 4H, Ar+H4thiaz), 4.02 (s, 2H, COCH <sub>2</sub> ), 3.13 (m, 1H, CHMe <sub>2</sub> ), 1.32 (d, 6H, CHMe <sub>2</sub> )	CDCl <sub>3</sub>
	170 - 172	12.05 (s broad, 1H, NHCO), 10.82 (s, 1H, NH), 7.48-6.90 (m, 5H, indole+H4thiaz), 3.74 (s, 2H, COCH <sub>2</sub> ), 3.06 (m, 1H, CHMe <sub>2</sub> ), 2.36 (s, 3H, Me), 1.21 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	163 - 165	12.07 (s broad, 1H, NHCO), 7.57-7.01 (m, 6H, indole+H4thiaz), 3.79 (s, 2H, COCH <sub>2</sub> ), 3.74 (s, 3H, NMe), 3.05 (m, 1H, CHMe <sub>2</sub> ), 1.21 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>

	155	10.20 (s broad, 1H, NHCO), 7.88-7.40 (m, 5H, Ar), 6.95 (s, 1H, H4thiaz), 4.04 (s, 2H, COCH <sub>2</sub> ), 3.07 (m, 1H, <u>CHMe</u> <sub>2</sub> ), 1.27 (d, 6H, CH <u>Me</u> <sub>2</sub> )	DMSO-d <sup>6</sup>
	-		
	157		
	234	11.3 (s broad, 1H, NHCO), 7.52-6.28 (m, 5H, Ar+H4thiaz), 3.93 (s, 2H, COCH <sub>2</sub> ), 3.87 (s, 3H, OMe), 3.10 (m, 1H, <u>CHMe</u> <sub>2</sub> ), 1.27 (d, 6H, CH <u>Me</u> <sub>2</sub> )	DMSO-d <sup>6</sup>
	236		
	161	12.19 (s, 1H, NHCO), 8.49-7.34 (m, 4H, Ar), 7.12 (s, 1H, H4thiaz), 2.56 (d, 2H, CH <sub>2</sub> iPr), 1.75 (m, 1H, <u>CHMe</u> <sub>2</sub> ), 0.86 (d, 6H, CH <u>Me</u> <sub>2</sub> )	DMSO-d <sup>6</sup>
	163		
	166	12.20 (s, 1H, NHCO), 8.48-7.24 (m, 10H, 2Xar+H4thiaz), 4.06 (s, 2H, CH <sub>2</sub> Ph), 3.77 (s, 2H, CH <sub>2</sub> CO)	DMSO-d <sup>6</sup>
	168		
	164	8.63-7.9 (m, 5H, Ar), 7.11 (s, 1H, H4thiaz), 3.85 (s, 2H, COCH <sub>2</sub> ), 3.15 (m, 1H, <u>CHMe</u> <sub>2</sub> ), 1.29 (d, 6H, CH <u>Me</u> <sub>2</sub> )	CDCl <sub>3</sub>
	167		
	114	11.6 (s broad, 1H, NHCO), 7.10 (s, 1H, H4thiaz), 3.67 (s, 3H, CH <sub>3</sub> OCO), 3.15 (m, 1H, <u>CHMe</u> <sub>2</sub> ), 2.60 (m, 2H, CH <sub>2</sub> CH <sub>2</sub> CH <sub>2</sub> ), 2.46 (m, 2H, CH <sub>2</sub> CH <sub>2</sub> CH <sub>2</sub> ), 2.09 (m, 2H, CH <sub>2</sub> CH <sub>2</sub> CH <sub>2</sub> ), 1.34 (d, 6H, CH <u>Me</u> <sub>2</sub> )	CDCl <sub>3</sub>
	117		
	217	10.6 (s broad, 1H, NHCO), 7.36 (m, 5H, Ar), 7.10 (s, 1H, H4thiaz), 6.61 (d, 1H, J=15.8, CH=CHPh), 6.36 (dt, 1H, J=15.8, 7.3, CH=CHPh), 3.43 (dd, 2H, J=7.3, 1.3, COCH <sub>2</sub> ), 3.14 (m, 1H, <u>CHMe</u> <sub>2</sub> ), 1.33 (d, 6H, CH <u>Me</u> <sub>2</sub> )	CDCl <sub>3</sub>
	220		
	222	12.09 (s broad, 1H, NHCO), 11.5 (s, 1H, NH), 7.78-7.16 (m, 4H, indole), 7.13 (s, 1H, H4thiaz), 3.78 (s, 2H, COCH <sub>2</sub> ), 3.07 (m, 1H, <u>CHMe</u> <sub>2</sub> ), 1.21 (d, 6H, CH <u>Me</u> <sub>2</sub> )	DMSO-d <sup>6</sup>
	225 dec .		
	222	12.07 (s, 1H, NHCO), 11.03 (s, 1H, NH), 7.3-6.80 (m, 5H, indole+H4thiaz), 3.77 (s, 2H, COCH <sub>2</sub> ), 3.06 (m, 1H, <u>CHMe</u> <sub>2</sub> ), 1.22 (d, 6H, CH <u>Me</u> <sub>2</sub> )	DMSO-d <sup>6</sup>
	225 dec .		

	172	12.25 (s, 1H, NHCO), 8.02-7.4 (m, 4H, Ar), 7.15 (s, 1H, H4thiaz), 4.0 (s, 2H, COCH <sub>2</sub> ), 3.07 (m, 1H, CHMe <sub>2</sub> ), 1.22 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	173	12.05 (s, 1H, NHCO), 10.77 (s, 1H, NH), 7.22-6.70 (m, 5H, indole+ H4thiaz), 3.75 (s, 2H, COCH <sub>2</sub> ), 3.72 (s, 3H, OMe), 3.07 (m, 1H, CHMe <sub>2</sub> ), 1.22 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	203	12.05 (s, 1H, NHCO), 10.77 (s, 1H, NH), 7.22-6.70 (m, 5H, indole+ H4thiaz), 3.75 (s, 2H, COCH <sub>2</sub> ), 3.72 (s, 3H, OMe), 3.07 (m, 1H, CHMe <sub>2</sub> ), 1.22 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	204	12.05 (s, 1H, NHCO), 10.77 (s, 1H, NH), 7.22-6.70 (m, 5H, indole+ H4thiaz), 3.75 (s, 2H, COCH <sub>2</sub> ), 3.72 (s, 3H, OMe), 3.07 (m, 1H, CHMe <sub>2</sub> ), 1.22 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	163	12.89 (s, 1H, NHCO), 10.75 (s, 1H, NH), 7.12-6.97 (m, 5H, indole+ H4thiaz), 3.10 (s, 1H, CHMe <sub>2</sub> ), 3.01 (t, 2H, CH <sub>2</sub> CH <sub>2</sub> CO), 2.78 (t, 2H, CH <sub>2</sub> CH <sub>2</sub> CO), 1.25 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	164	12.89 (s, 1H, NHCO), 10.75 (s, 1H, NH), 7.12-6.97 (m, 5H, indole+ H4thiaz), 3.10 (s, 1H, CHMe <sub>2</sub> ), 3.01 (t, 2H, CH <sub>2</sub> CH <sub>2</sub> CO), 2.78 (t, 2H, CH <sub>2</sub> CH <sub>2</sub> CO), 1.25 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	186	12.7 (s broad, 1H, NHCO), 8.18 (d, 1H, J=7.8, Ar), 7.71 (d, 1H, J=7.8, Ar), 7.24 (s, 1H, H4thiaz), 3.15 (m, 1H, CHMe <sub>2</sub> ), 1.27 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	187	12.7 (s broad, 1H, NHCO), 8.18 (d, 1H, J=7.8, Ar), 7.71 (d, 1H, J=7.8, Ar), 7.24 (s, 1H, H4thiaz), 3.15 (m, 1H, CHMe <sub>2</sub> ), 1.27 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	108	10.8 (s broad, 1H, NHCO), 7.45 (s, 1H, H4thiaz), 3.33 (m, 1H, CHMe <sub>2</sub> ), 2.54 (m, 2H, CH <sub>2</sub> CHMe <sub>2</sub> ), 2.42 (m, 1H, CH <sub>2</sub> CHMe <sub>2</sub> ), 1.53 (d, 6H, CH <sub>2</sub> CHMe <sub>2</sub> ), 1.21 (d, 6H, CHMe <sub>2</sub> )	CDCl <sub>3</sub>
	124	12.4 (s broad, 1H, NHCO), 8.05-7.51 (m, 5H, Ph), 7.23 (s, 1H, H4thiaz), 3.13 (m, 1H, CHMe <sub>2</sub> ), 1.28 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	118	11.8 (s broad, 1H, NHCO), 7.11 (s, 1H, H4thiaz), 3.08 (m, 1H, CHMe <sub>2</sub> ), 2.25 (d, 2H, CH <sub>2</sub> CO), 2.42 (m, 1H, CH <sub>2</sub> CHMe <sub>2</sub> ), 1.23 (d, 6H, CHMe <sub>2</sub> ), 1.8-0.8 (m, 11H, cyclohexyl)	DMSO-d <sup>6</sup>
	132	8.13 (d, 1H, H3fur), 7.84 (d, 1H, H5fur), 7.25 (d, 1H, H4thiaz), 6.69 (dd, 1H, H4fur), 7.45 (s, 1H, H4thiaz), 3.20 (m, 1H, CHMe <sub>2</sub> ), 1.39 (d, 6H, CHMe <sub>2</sub> )	CDCl <sub>3</sub>
	133	12.7 (s broad, 1H, NHCO), 7.54-6.82 (m, 3H, H4thiaz+furane), 3.10 (m, 1H, CHMe <sub>2</sub> ), 1.26 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>
	134	12.7 (s broad, 1H, NHCO), 7.54-6.82 (m, 3H, H4thiaz+furane), 3.10 (m, 1H, CHMe <sub>2</sub> ), 1.26 (d, 6H, CHMe <sub>2</sub> )	DMSO-d <sup>6</sup>

## CLAIMS

1. The use of a compound which is a 2-amino-1,3-thiazole derivative of formula (I)



5 wherein

R is a halogen atom, a nitro group, an optionally substituted amino group or it is a group, optionally further substituted, selected from:

i) straight or branched C<sub>1</sub>-C<sub>8</sub> alkyl, C<sub>2</sub>-C<sub>6</sub> alkenyl or C<sub>2</sub>-C<sub>6</sub> alkynyl;

10 ii) C<sub>3</sub>-C<sub>6</sub> cycloalkyl;

iii) aryl or arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

R<sub>1</sub> is an optionally further substituted group selected from:

i) straight or branched C<sub>1</sub>-C<sub>8</sub> alkyl or C<sub>2</sub>-C<sub>6</sub> alkenyl;

15 ii) 3 to 6 membered carbocycle or 5 to 7 membered heterocycle ring;

iii) aryl or arylcarbonyl;

20 iv) arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

v) arylalkenyl with from 2 to 6 carbon atoms within the straight or branched alkenyl chain;

25 vi) an optionally protected amino acid residue;

or a pharmaceutically acceptable salt thereof; in the manufacture of a medicament for treating cell proliferative disorders associated with an altered cell dependent kinase activity.

30 2. Use according to claim 1 wherein the said cell proliferative disorder is selected from the group consisting of cancer, Alzheimer's disease, viral infections, auto-immune diseases or neurodegenerative disorders.

3. Use according to claim 2 wherein the cancer is selected from the group consisting of carcinoma, squamous cell carcinoma, hematopoietic tumors of myeloid or lymphoid lineage, tumors of mesenchymal origin, tumors of the 5 central and peripheral nervous system, melanoma, seminoma, teratocarcinoma, osteosarcoma, xenoderoma pigmentosum, keratoctanthoma, thyroid follicular cancer and Kaposi's sarcoma.

10 4. Use according to claim 1 wherein the cell proliferative disorder is selected from the group consisting of benign prostate hyperplasia, familial adenomatosis polyposis, neuro-fibromatosis, psoriasis, vascular smooth cell proliferation associated with 15 atherosclerosis, pulmonary fibrosis, arthritis glomerulonephritis and post-surgical stenosis and restenosis.

20 5. Use according to any one of the preceding claims wherein the medicament enables tumor angiogenesis and metastasis inhibition.

25 6. A compound which is a 2-amino-1,3-thiazole derivative of formula (I)

wherein

R is a halogen atom, a nitro group, an optionally substituted amino group or it is a group, optionally further substituted, selected from:

30 i) straight or branched C<sub>1</sub>-C<sub>8</sub> alkyl, C<sub>2</sub>-C<sub>6</sub> alkenyl or C<sub>2</sub>-C<sub>6</sub> alkynyl;

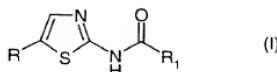
ii) C<sub>3</sub>-C<sub>6</sub> cycloalkyl;

iii) aryl or arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

$R_1$  is an optionally further substituted group selected from:

- i) straight or branched  $C_1$ - $C_8$  alkyl or  $C_2$ - $C_6$  alkenyl;
- ii) 3 to 6 membered carbocycle or 5 to 7 membered heterocycle ring;
- 5      iii) aryl or arylcarbonyl;
- iv) arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;
- v) arylalkenyl with from 2 to 6 carbon atoms within the straight or branched alkenyl chain;
- 10     vi) an optionally protected amino acid residue; or a pharmaceutically acceptable salt thereof; for use as a medicament; provided that each of  $R$  and  $R_1$ , independently, is not a methyl group and that the compound is not 2-diethylaminomethyl-carbonylamino-5-chloro-1,3-thiazole.
- 15

7. A compound which is a 2-amino-1,3-thiazole derivative of formula (I)



20    wherein

$R$  is a halogen atom, a nitro group, an optionally substituted amino group or it is a group, optionally further substituted, selected from:

- i) straight or branched  $C_1$ - $C_8$  alkyl,  $C_2$ - $C_6$  alkenyl or  $C_2$ - $C_6$  alkynyl;
- 25    ii)  $C_3$ - $C_6$  cycloalkyl;
- iii) aryl or arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

$R_1$  is an optionally further substituted group selected from:

- i) straight or branched  $C_1$ - $C_8$  alkyl or  $C_2$ - $C_6$  alkenyl;
- ii) 3 to 6 membered carbocycle or 5 to 7 membered heterocycle ring;
- 35    iii) aryl or arylcarbonyl;
- iv) arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

- v) arylalkenyl with from 2 to 6 carbon atoms within the straight or branched alkenyl chain;
- vi) an optionally protected amino acid residue; or a pharmaceutically acceptable salt thereof;

5 provided that:

- a) R and R<sub>1</sub>, each independently, are not methyl;
- b) when R is bromine or chlorine then, R<sub>1</sub> is not unsubstituted C<sub>1</sub>-C<sub>4</sub> alkyl or an optionally substituted aminomethyl;

10 c) when R is nitro or phenyl, then R<sub>1</sub> is not unsubstituted phenyl.

8. A compound of formula (I), according to claim 7, wherein R is a halogen atom or an optionally substituted group selected from a straight or branched C<sub>1</sub>-C<sub>4</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> cycloalkyl, aryl or an arylalkyl with from 1 to 4 carbon atoms within the alkyl chain; R<sub>1</sub> is an optionally substituted group selected from straight or branched C<sub>1</sub>-C<sub>4</sub> alkyl or alkenyl, aryl or arylalkyl with from 1 to 4 carbon atoms within the alkyl chain or it is an optionally protected amino acid residue.

9. A compound of formula (I), according to claim 8, wherein R is a bromine or chlorine atom or is an optionally substituted group selected from straight or branched C<sub>1</sub>-C<sub>4</sub> alkyl, cyclopropyl, aryl or arylalkyl with from 1 to 2 carbon atoms within the alkyl chain; R<sub>1</sub> is an optionally substituted group selected from straight or branched C<sub>1</sub>-C<sub>4</sub> alkyl or alkenyl, aryl or arylalkyl with from 1 to 4 carbon atoms within the alkyl chain or it is an optionally protected amino acid residue.

10. A compound of formula (I) according to claim 7 wherein R is a halogen atom or is selected from nitro, amino, alkylamino, hydroxyalkylamino, arylamino, C<sub>3</sub>-C<sub>6</sub> cycloalkyl and straight or branched C<sub>1</sub>-C<sub>6</sub> alkyl which is unsubstituted or substituted by hydroxy, alkylthio,

alkoxy, amino, alkylamino, alkoxy carbonylamino, alkoxy carbonyl alkylamino, alkyl carbonyl, alkylsulfonyl, alkoxy carbonyl, carboxy or aryl which is unsubstituted or substituted by one or more hydroxy, halogen, nitro, 5 alkoxy, aryloxy, alkylthio, arylthio, amino, alkylamino, dialkylamino, N-alkyl-piperazinyl, 4-morpholinyl, arylamino, cyano, alkyl, phenyl, aminosulfonyl, aminocarbonyl, alkylcarbonyl, arylcarbonyl, alkoxy carbonyl or carboxy groups, or R is 10 an aryl group which is unsubstituted or substituted by one or more hydroxy, halogen, nitro, alkoxy, aryloxy, alkylthio, arylthio, amino, alkylamino, dialkylamino, N-alkyl-piperazinyl, 4-morpholinyl, arylamino, cyano, alkyl, phenyl, aminosulphonyl, aminocarbonyl, 15 alkylcarbonyl, arylcarbonyl, alkoxy carbonyl or carboxy groups;

R<sub>1</sub> is a straight or branched C<sub>1</sub>-C<sub>6</sub> alkyl group or an aryl group, each being unsubstituted or substituted as defined above for R;

20 or a pharmaceutically acceptable salt thereof;  
provided that:

- a) R and R<sub>1</sub>, each independently, are not methyl;
- b) when R is bromine or chlorine then, R<sub>1</sub> is not unsubstituted C<sub>2</sub>-C<sub>6</sub> alkyl or an optionally substituted 25 aminomethyl;
- c) when R is nitro or phenyl, then R<sub>1</sub> is not unsubstituted phenyl.

30 11. A compound of formula (I) according to any one of the preceding claims, whenever appropriate in the form of a pharmaceutically acceptable salt, selected from the group consisting of:

1. ethyl 3-[(5-bromo-1,3-thiazol-2-yl)amino]-3-oxopropanoate;
- 35 2. N-(5-bromo-1,3-thiazol-2-yl)-2-phenyl-acetamide;
3. N-(5-bromo-1,3-thiazol-2-yl)-benzamide;

4. Ethyl 4-[(5-bromo-1,3-thiazol-2-yl)amino]-4-oxobutanoate;

5. N-(5-Bromo-thiazol-2-yl)-3-hydroxy-propionamide;

6. N-(5-Bromo-1,3-thiazol-2-yl)-4-hydroxybutanamide;

7. N-(5-Bromo-thiazol-2-yl)-2-ethoxy-acetamide;

5 8. 2-N-[2-(3-pyridyl)-acetyl-amino]-5-bromo-thiazole;

9. 2-N-[2-(3-pyridyl)-acetyl-amino]-5-isopropyl-thiazole;

10. N-(5-bromo-1,3-thiazol-2-yl)-2-(3-hydroxyphenyl)acetamide;

11. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-hydroxyphenyl)acetamide;

10 12. N-(5-bromo-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide;

13. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide;

15 14. N-(5-bromo-1,3-thiazol-2-yl)-2-(3-chorophenyl)acetamide;

15. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-chorophenyl)acetamide;

16. N-(5-bromo-1,3-thiazol-2-yl)-2-(4-hydroxyphenyl)acetamide;

20 17. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxyphenyl)acetamide;

18. N-(5-bromo-1,3-thiazol-2-yl)-2-(3,4-dihydroxyphenyl)acetamide;

19. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dihydroxyphenyl)acetamide;

25 20. N-(5-bromo-1,3-thiazol-2-yl)-2-(4-hydroxy-3-methoxyphenyl)acetamide;

21. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxy-3-methoxyphenyl)acetamide;

30 22. N-(5-bromo-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide;

23. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide;

24. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-chlorophenyl)acetamide;

35 25. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-acetamide;

26. N-(5-bromo-thiazol-2-yl)-4-sulfamoyl-benzamide;

27. N-(5-isopropyl-thiazol-2-yl)-4-sulfamoyl-benzamide;  
28. 4-amino-N-(5-bromo-1,3-thiazol-2-yl)butanamide;  
29. 3-amino-N-(5-bromo-1,3-thiazol-2-yl)propionamide;  
30. N-(5-isopropyl-1,3-thiazol-2-yl)-butanamide;  
5 31. N-(5-bromo-1,3-thiazol-2-yl)-butanamide;  
32. N-(5-chloro-1,3-thiazol-2-yl)-butanamide;  
33. N-(5-phenyl-1,3-thiazol-2-yl)-butanamide;  
34. N-(5-nitro-1,3-thiazol-2-yl)-butanamide;  
35. N-(5-methyl-1,3-thiazol-2-yl)-butanamide;  
10 36. N-(5-benzyl-1,3-thiazol-2-yl)-butanamide;  
37. N-(5-isobutyl-1,3-thiazol-2-yl)-butanamide;  
38. N-(5-cyclopropyl-1,3-thiazol-2-yl)-butanamide;  
39. N-[5-[2-(methylsulfonyl)ethyl]-1,3-thiazol-2-yl]-  
butanamide;  
15 40. N-[5-(2-methylthioethyl)-1,3-thiazol-2-yl]-butanamide;  
41. N-[5-[2-(methoxycarbonyl)ethyl]-1,3-thiazol-2-yl]-  
butanamide;  
42. N-[5-(3-methoxy-propyl)-1,3-thiazol-2-yl]-butanamide;  
43. N-[5-(2-ethoxy-ethyl)-1,3-thiazol-2-yl]-butanamide;  
20 44. N-[5-(indol-3-yl-methyl)-1,3-thiazol-2-yl]-butanamide;  
45. N-[5-(3-oxo-butyl)-1,3-thiazol-2-yl]-butanamide;  
46. 2-[3-(3-chloropropoxy)phenyl]-N-(5-isopropyl-1,3-  
thiazol-2-yl)acetamide;  
47. 2-[3-(2-chloroethoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-  
25 2-yl)acetamide;  
48. 2-(4-aminophenyl)-N-(5-isopropyl-1,3-thiazol-2-  
yl)acetamide;  
49. 4-amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;  
50. 2-(2-amino-1,3-thiazol-4-yl)-N-(5-isopropyl-1,3-thiazol-  
30 2-yl)acetamide;  
51. N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[3-(4-  
morpholinyl)propoxy]phenyl}acetamide;  
52. N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[2-(4-  
morpholinyl)ethoxy]phenyl}acetamide;  
35 53. N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[3-(1-  
pirrolidinyl)propoxy]phenyl}acetamide;

54. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-[3-(4-methyl-1-piperazinyl)propoxy]phenyl)acetamide;

55. 2-[3-[2-(dimethylamino)ethoxy]phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;

5 56. 2-(3-[3-(dimethylamino)propoxy]phenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

57. 2-[4-(dimethylamino)phenyl]-N-(5-isobutyl-1,3-thiazol-2-yl)acetamide

58. 2-(1,3-benzodioxol-5-yl)-N-(5-isobutyl-1,3-thiazol-2-yl)acetamide

10 59. N-(5-benzyl-1,3-thiazol-2-yl)-2-[4-(dimethylamino)phenyl]acetamide

60. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(2-methoxyethoxy)-phenyl]acetamide

15 61. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-methyl-1-piperazinyl)benzamide

62. N-(5-isobutyl-1,3-thiazol-2-yl)-2-(3-pyridinyl)acetamide

63. N-(5-benzyl-1,3-thiazol-2-yl)-2-(3-pyridinyl)acetamide

64. 2-[N-[2'-N'-(ethoxycarbonyl-methyl)-amino]-acetyl]-20 amino-5-bromo-thiazole

65. 2-anilino-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

66. (R)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide

67. (S)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide

68. N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

25 69. 2,5-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

70. 3,5-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

71. 3,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

72. 2,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

73. 2,3-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

30 74. 3-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

75. 2-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

76. 4-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

77. 3-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

78. 4-chloro-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-35 yl)benzamide

79. 5-bromo-2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

80. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

81. 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
82. 4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
83. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
84. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
5 85. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
86. 2,4-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
87. 3,4-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
88. 2,3,4,5,6-pentafluoro-N-(5-isopropyl-1,3-thiazol-2-  
y1)benzamide  
10 89. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-3-  
nitrobenzamide  
90. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-  
nitrobenzamide  
91. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-  
15 nitrobenzamide  
92. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-  
nitrobenzamide  
93. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxy-2-  
nitrobenzamide  
20 94. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-2-  
nitrobenzamide  
95. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxy-3-  
nitrobenzamide  
96. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-  
25 nitrobenzamide  
97. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dinitrobenzamide  
98. 5-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl)-2-  
nitrophenyl octanoate  
99. N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide  
30 100. N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide  
101. N-(5-isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamide  
102. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl)-3-  
nitrobenzamide  
103. 4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-  
35 nitrobenzamide  
104. 6-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-  
nitrobenzamide

105. 4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide

106. 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamide

5 107. 5-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide

108. 2-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-5-nitrobenzamide

109. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide

110. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide

111. N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitro-4-(trifluoromethyl)benzamide

15 112. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-bis(trifluoromethyl)benzamide

113. N-(5-isopropyl-1,3-thiazol-2-yl)-2,6-bis(trifluoromethyl)benzamide

114. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(trifluoromethyl)benzamide

20 115. N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide

116. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethyl)benzamide

25 117. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide

118. 5-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide

119. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethyl)benzamide

30 120. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide

121. methyl 4-{{(5-isopropyl-1,3-thiazol-2-yl)amino}carbonyl}benzoate

35 122. methyl 2-{{(5-isopropyl-1,3-thiazol-2-yl)amino}carbonyl}benzoate

123. 4-cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

124. 3-cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

125. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methylbenzamide  
126. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methylbenzamide  
127. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylbenzamide  
128. N-(5-isopropyl-1,3-thiazol-2-yl)-4-vinylbenzamide  
5 129. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(2-phenylethynyl)benzamide  
130. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-methylbenzamide  
131. 2-benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
10 132. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenethylbenzamide  
133. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylbenzamide  
134. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenylbenzamide  
135. 4-(tert-butyl)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
15 136. N-(5-isopropyl-1,3-thiazol-2-yl)-4-isopropylbenzamide  
137. N-(5-isopropyl-1,3-thiazol-2-yl)-4-pentylbenzamide  
138. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylbenzamide  
139. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4-dimethylbenzamide  
20 140. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethylbenzamide  
141. 4-acetyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
142. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl)benzamide  
143. 5-(aminosulfonyl)-2,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
25 144. 5-(aminosulfonyl)-4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide  
145. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide  
30 146. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide  
147. 5-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxybenzamide  
148. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide  
35 149. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxybenzamide  
150. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxybenzamide  
151. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4-dimethoxybenzamide

152. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethoxybenzamide

153. N-(5-isopropyl-1,3-thiazol-2-yl)-2,4-dimethoxybenzamide

5 154. N-(5-isopropyl-1,3-thiazol-2-yl)-2,3-dimethoxybenzamide

155. N-(5-isopropyl-1,3-thiazol-2-yl)-3-phenoxybenzamide

156. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenoxybenzamide

157. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenoxybenzamide

10 158. 2-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

159. 4-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

160. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4,5-trimethoxybenzamide

161. 3,4-diethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

15 162. 3,4,5-triethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

163. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-(methoxymethoxy)benzamide

164. 4-butoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

20 165. N-(5-isopropyl-1,3-thiazol-2-yl)-4-propoxybenzamide

166. 4-isopropoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

167. N-(5-isopropyl-1,3-thiazol-2-yl)-1,3-benzodioxole-5-carboxamide

168. 4-(benzyloxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

25 169. 4-(2-cyclohexen-1-yloxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

170. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethoxy)benzamide

30 171. 4-(difluoromethoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

172. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfanyl)benzamide

173. 2-[(4-chlorophenyl)sulfinyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

35 174. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[(4-nitrophenyl)sulfinyl]benzamide

175. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(4-methylphenyl)sulfonyl]-3-nitrobenzamide

176. N-(5-isopropyl-1,3-thiazol-2-yl)-3-[(trifluoromethyl)sulfanyl]benzamide

5 177. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxy-4-(methylsulfanyl)benzamide

178. 2-[(2-cyanophenyl)sulfanyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

179. N-1~,N-1~-diethyl-3,6-difluoro-N-2--(5-isopropyl-1,3-thiazol-2-yl)phthalamide

10 180. 4-formyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

181. 2-formyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

182. 4-{{(2,5-dimethoxyanilino)carbonyl}amino}-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

15 183. 4-(hydroxymethyl)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

184. 4-{{(5-isopropyl-1,3-thiazol-2-yl)amino}carbonyl}-2-nitrobenzyl acetate

185. 4-{{(5-isopropyl-1,3-thiazol-2-yl)amino}carbonyl}-2-nitrobenzyl 4-(acetylamino)-3-iodobenzoate

20 186. 4-(acetylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

187. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(2-phenylacetyl)amino]benzamide

25 188. 4-(acetylamino)-3-iodo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

189. 4-amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

190. 4-(dimethylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

30 191. 3-(dimethylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

192. 2-(methylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

193. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(trifluoromethyl)anilino]benzamide

35 194. 3-{{(5-bromo-1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)methyl}amino}-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

195. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(1H-pyrrol-1-yl)benzamide

196. 2,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)isonicotinamide

5 197. 2-(4-bromophenyl)-6-(4-iodophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)isonicotinamide

198. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(trifluoromethyl)anilino]nicotinamide

199. 2,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide

10 200. 5,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide

201. 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-6-methylnicotinamide

15 202. 2,6-dichloro-5-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide

203. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenoxy nicotinamide

204. N-(5-isopropyl-1,3-thiazol-2-yl)-6-(2,2,2-trifluoroethoxy)nicotinamide

20 205. N-(5-isopropyl-1,3-thiazol-2-yl)-2,6-dimethoxy nicotinamide

206. N-(5-isopropyl-1,3-thiazol-2-yl)-2-quinoxalinecarboxamide

207. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-pyrazinecarboxamide

25 208. N-(5-isopropyl-1,3-thiazol-2-yl)-8-quinolinecarboxamide

209. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-4-quinolinecarboxamide

30 210. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-1-phenyl-1H-pyrazole-4-carboxamide

211. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-1H-pyrazole-3-carboxamide

212. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-pyrazole-4-carboxamide

35 213. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-phenyl-2H-1,2,3-triazole-4-carboxamide

214. 2-[(2,1,3-benzoxadiazol-5-yloxy)methyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-1,3-thiazole-5-carboxamide

215. N-(5-isopropyl-1,3-thiazol-2-yl)-9H-fluorene-1-carboxamide

216. N-(5-isopropyl-1,3-thiazol-2-yl)-7-methoxy-1-benzofuran-2-carboxamide

217. N-(5-isopropyl-1,3-thiazol-2-yl)-1-[(4-methylphenyl)sulfonyl]-1H-pyrrole-3-carboxamide

218. 2-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamide

219. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamide

220. N-(5-isopropyl-1,3-thiazol-2-yl)-2-naphthamide

221. N-(5-isopropyl-1,3-thiazol-2-yl)-9,10-dioxo-9,10-dihydro-2-anthracenecarboxamide

222. N-(5-isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluorene-4-carboxamide

223. N-(5-isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluorene-1-carboxamide

224. N-(5-isopropyl-1,3-thiazol-2-yl)-8-oxo-5,6,7,8-tetrahydro-2-naphthalenecarboxamide

225. N-(5-isopropyl-1,3-thiazol-2-yl)-1,3-dioxo-1,3-dihydro-2-benzofuran-5-carboxamide

226. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-5-carboxamide

227. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-4-carboxamide

228. N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-2-phenyl-1H-indole-5-carboxamide

229. 2-butyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indole-5-carboxamide

230. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-6-carboxamide

231. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methoxy-1H-indole-2-carboxamide

232. 1-allyl-2-butyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-5-carboxamide

233. N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indole-2-carboxamide

234. 1-benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-1H-indole-5-carboxamide

5 235. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-1,2,3-benzotriazole-5-carboxamide

236. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-isoxazolecarboxamide

10 237. N-(5-isopropyl-1,3-thiazol-2-yl)-3-thiophenecarboxamide

238. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-thiophenecarboxamide

239. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-thiophenecarboxamide

15 240. 5-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-2-thiophenecarboxamide

241. N-(5-isopropyl-1,3-thiazol-2-yl)-3-[(2,3,3-trichloroacryloyl)amino]-2-thiophenecarboxamide

242. 5-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide

20 243. N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide

244. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(4-nitrophenyl)-2-furamide

245. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(2-nitrophenyl)-2-furamide

25 246. 5-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide

247. N-(5-isopropyl-1,3-thiazol-2-yl)-5-[3-(trifluoromethyl)phenyl]-2-furamide

248. 5-(4-chloro-2-nitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide

30 249. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(4-methyl-2-nitrophenyl)-2-furamide

250. 5-[2-chloro-5-(trifluoromethyl)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide

35 251. tert-butyl (1R)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethylcarbamate

252. (1R)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate

253. (1S)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate

254. (R,S)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

5 255. (R)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

256. (S)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

10 257. 2-(acetylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

258. (R,S)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

15 259. (R)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

260. (S)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

261. 3,3,3-trifluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxy-2-phenylpropanamide

262. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(1-

20 naphthyl)acetamide

263. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-naphthyl)acetamide

264. 2-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

25 265. 2-(1,3-benzodioxol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

266. 2-(2,4-dinitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

267. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-methyl-1H-indol-3-yl)acetamide

30 268. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(1-methyl-1H-indol-3-yl)acetamide

269. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(5-methoxy-1H-indol-3-yl)acetamide

35 270. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(5-benzyloxy-1H-indol-3-yl)acetamide

271. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxy-2-methyl-1H-indol-3-yl)acetamide

272. 2-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxoacetamide

273. 2-(5-bromo-1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

5 274. 2-(5-fluoro-1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

275. 2-[1-(4-chlorobenzoyl)-5-methoxy-2-methyl-1H-indol-3-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

276. 3-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

10 277. 4-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)butanamide

278. N-(5-isopropyl-1,3-thiazol-2-yl)-3-(2-thienyl)propanamide

15 279. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-thienyl)acetamide

280. N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxo-2-(2-thienyl)acetamide

281. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-thienyl)acetamide

20 282. 2-(5-chloro-1-benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

283. 2-(1-benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

25 284. 2-[2-(formylamino)-1,3-thiazol-4-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamide

285. 2-{2-[(2-chloroacetyl)amino]-1,3-thiazol-4-yl}-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamide

286. 2-chloro-N-(4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}-1,3-thiazol-2-yl)acetamide

30 287. ethyl 2-({[2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-(1H-pyrazol-3-yl)ethylidene]amino}oxy)acetate

288. 2-(2-furyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxoacetamide

35 289. 2-(5-bromo-3-pyridinyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

290. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(7-methoxy-2-oxo-2H-chromen-4-yl)acetamide

291. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenyl-3-butenamide  
292. N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxo-4-(4-methyl-  
phenyl)butanamide  
293. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-  
5 nitrophenyl)butanamide  
294. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenylbutanamide  
295. benzyl 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]-4-  
oxobutylcarbamate  
296. methyl 5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-  
10 oxopentanoate  
297. 4-(1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)-N-(5-  
isopropyl-1,3-thiazol-2-yl)butanamide  
298. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-methoxy-1-  
naphthyl)-4-oxobutanamide  
299. 3-(2-chlorophenoxy)-N-(5-isopropyl-1,3-thiazol-2-  
15 yl)propanamide  
300. 3-(4-methylphenoxy)-N-(5-isopropyl-1,3-thiazol-2-  
yl)propanamide  
301. 3-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-  
20 yl)propanamide  
302. 3-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-  
yl)propanamide  
303. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylpentanamide  
304. 3-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-  
25 yl)propanamide  
305. 3-(4-methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-  
yl)propanamide  
306. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide  
307. 3-phenyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide  
308. 2-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide  
309. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methylbutanamide  
310. N-(5-isopropyl-1,3-thiazol-2-yl)-5-oxo-5-  
phenylpentanamide  
311. 2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-  
35 phenylethyl acetate  
312. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[4-(1-oxo-1,3-  
dihydro-2H-isoindol-2-yl)phenyl]propanamide

313. 1-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)cyclopentanecarboxamide

314. 1-phenyl-N-(5-isopropyl-1,3-thiazol-2-yl)cyclopentanecarboxamide

5 315. 2-(3-bromo-4-methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

316. 2-(2-nitro-4-trifluoromethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

317. 5-cyclohexyl 1-(4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}benzyl) (2S)-2-[(tert-butoxycarbonyl)amino]pentanedioate

10 318. 2-(5,6-dimethyl-1H-benzimidazol-1-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

319. 2-[5-(4-chlorophenyl)-2H-1,2,3,4-tetraazol-2-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

15 320. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[5-(1-pyrrolidinyl)-2H-1,2,3,4-tetraazol-2-yl]acetamide

321. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methyl-1-benzothiophen-2-yl)acetamide

20 322. N-(5-isopropyl-1,3-thiazol-2-yl)-4,4-bis(4-methylphenyl)-3-butenamide

323. 2-cyclopropyl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

324. N-(4-bromo-6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexyl)benzamide

25 325. 2-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

326. benzyl 6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexylcarbamate

30 327. N-1-(5-isopropyl-1,3-thiazol-2-yl)-N-4-(2-propynyl)-2-butenediamide

328. 4-(2,4-dimethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide

329. 4-(4-benzyloxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide

35 330. 4-(thiophen-2-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide

331. benzyl 2-[(benzyloxy)carbonyl]amino]-5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate

332. 4-(1H-indol-3-yl)-N-{3-[(5-isopropyl-1,3-thiazol-2-yl)amino]-3-oxopropyl}butanamide

5 333. 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonylphenyl 4-chlorobenzenesulfonate

334. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(2-methoxyanilino)carbonyl]amino)benzamide

10 335. 4-[(2-isopropylsulfonyl)acetyl]amino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

336. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(2-phenylsulfanyl)acetyl]amino)benzamide

337. 4-[(diethylamino)sulfonyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

15 338. 2-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

339. 3,5-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

340. 3-[(2-fluoroanilino)carbonyl]amino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

341. N-(5-isopropyl-1,3-thiazol-2-yl)-1-phenyl-5-propyl-1H-pyrazole-4-carboxamide

20 342. 3-chloro-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfanyl)-2-thiophenecarboxamide

343. 3-iodo-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfanyl)-2-thiophenecarboxamide

25 344. 2-[(4-chlorophenyl)sulfonylmethyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-1,3-thiazole-5-carboxamide

345. 5-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(trifluoromethyl)-3-furamide

30 346. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,3,4,5,6-pentafluorophenyl)acetamide

347. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-fluorophenyl)acetamide

35 348. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-bromophenyl)acetamide

349. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-chlorophenyl)acetamide

350. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-nitrophenyl)acetamide

351. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-trifluoromethylphenyl)acetamide

5 352. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-methoxyphenyl)acetamide

353. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-dimethoxyphenyl)acetamide

354. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-difluorophenyl)acetamide

10 355. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4,5-trimethoxyphenyl)acetamide

356. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,6-dichlorophenyl)acetamide

15 357. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-chloro-6-fluorophenyl)acetamide

358. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,5-dimethoxyphenyl)acetamide

359. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,5-difluorophenyl)acetamide

20 360. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-bis-trifluoromethylphenyl)acetamide

361. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methylthiophenyl)acetamide

25 362. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide

363. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-bromophenyl)acetamide

364. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-chlorophenyl)acetamide

30 365. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-fluorophenyl)acetamide

366. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl)acetamide

35 367. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-trifluoromethylphenyl)acetamide

368. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methylphenyl)acetamide

369. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-dimethylaminophenyl)acetamide

370. 2-[1,1'-biphenyl]-4-yl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

5 371. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-trifluoromethylphenyl)acetamide

372. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-bromophenyl)acetamide

373. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-chlorophenyl)acetamide

10 374. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-nitrophenyl)acetamide

375. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide

15 376. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-dinitrophenyl)acetamide

377. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-dichlorophenyl)acetamide

378. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-difluorophenyl)acetamide

20 379. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-benzyloxy-3-methoxyphenyl)acetamide

380. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dichlorophenyl)acetamide

25 381. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-difluorophenyl)acetamide

382. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dimethoxyphenyl)acetamide

383. 2-(2,3-dihydro-1H-inden-5-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

30 384. N-(5-isopropyl-1,3-thiazol-2-yl)-1-phenylcyclopropanecarboxamide

385. 2-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1-phenylacetamide

35 386. 2-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide

387. N-(5-isopropyl-1,3-thiazol-2-yl)-2,2-diphenylacetamide

388. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-nitrophenoxy)acetamide

389. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl)propanamide

5 390. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide

391. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-isobutylphenyl)propanamide

392. N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxo-2-phenylacetamide

10 393. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-phenylpentanamide

394. (E, Z)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-2-butenamide

395. N-(5-isopropyl-1,3-thiazol-2-yl)bicyclo[4.2.0]octa-1,3,5-triene-7-carboxamide

15 396. N-(5-isopropyl-1,3-thiazol-2-yl)-3-oxo-1-indanecarboxamide

397. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylbutanamide

398. tert-butyl (1S)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-1-methyl-2-oxoethylcarbamate

20 399. tert-butyl (1S,2S)-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl-2-methylbutylcarbamate

400. tert-butyl 2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

25 401. tert-butyl (1S)-5-amino-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonylpentylcarbamate

402. tert-butyl 4-[(imino[(4-methylphenyl)sulfonyl]amino)methyl]amino]-1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonylbutylcarbamate

30 403. tert-butyl 1-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl-3-(tritylamo)propylcarbamate

404. tert-butyl (1S)-1-(benzyloxymethyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

35 405. tert-butyl (1S)-1-benzyl-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

406. tert-butyl (1R)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-(benzylthiomethyl)ethylcarbamate

407. benzyl (3S)-3-[(tert-butoxycarbonyl)amino]-4-[(5-isopropyl-1,3-thiazol-2-yl)amino]-4-oxobutanoate

408. tert-butyl (2S)-2-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-1-pyrrolidinecarboxylate

5 409. tert-butyl (1S)-1-(1H-indol-3-ylmethyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

410. tert-butyl (1S)-1-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-3-(methylsulfanyl)propylcarbamate

411. tert-butyl (1S)-2-benzyloxy-1-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}propylcarbamate

10 412. tert-butyl (1S)-1-(4-benzyloxybenzyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

413. tert-butyl (1S)-1-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-2-methylpropylcarbamate

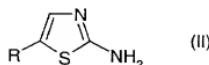
15 414. tert-butyl (1S)-1-{[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}-3-methylbutylcarbamate

415. benzyl (4S)-4-[(tert-butoxycarbonyl)amino]-5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate

and the pharmaceutically acceptable salts thereof.

20

12. A process for producing a compound of formula (I), as defined in claim 7, which process comprises reacting a compound of formula (II)



25 with a compound of formula (IV)



wherein R and  $R_1$  are as defined in claim 7 and X is hydroxy or a suitable leaving group;

and, if desired, converting a 2-amino-1,3-thiazole derivative of formula (I) into another such derivative of formula (I), and/or into a salt thereof.

13. A process according to claim 12 wherein X is hydroxy, bromine or chlorine.

**14.** A pharmaceutical composition comprising one or more pharmaceutically acceptable carriers and/or diluents and, as the active principle, an effective amount of a compound of formula (I) as defined in claim 7.

# Declaration and Power of Attorney for Patent Application

## Dichiarazione e procura ai fini della domanda di brevetto

### Italian Language Declaration

Il sottoscritto inventore dichiara che:

As a below named inventor, I hereby declare that:

La propria residenza, recapito postale e cittadinanza corrispondono a quanto indicato in calce, sotto la propria firma.

My residence, post office address and citizenship are as stated next to my name

Ritiene di essere il primo ed unico inventore originale (se viene elencato in calce un solo nominativo) o il coinventore primo ed originale (se è elencato più di un nominativo) del oggetto rivendicato e per il quale il sottoscritto presenta domanda di brevetto. La invenzione in questione è chiamata

I believe I am the original, first and sole inventor (if only one name is listed below) or an original, first and joint inventor (if plural names are listed below) of the subject matter which is claimed and for which a patent is sought on the invention entitled

2-Amino-thiazole derivatives, process  
for their preparation, and their use as  
antitumor agents.

e la sua descrizione è allegata alla presente Dichiarazione a meno

the specification of which

è qui allegato

is attached hereto

II \_\_\_\_\_

was filed on 27 October 1999

è stata depositata una domanda di brevetto statunitense numero o una domanda di brevetto internazionale PCT numero

as United States Application Number or PCT International Application Number

PCT/EP99/08306

\_\_\_\_\_ che è stata modificata il

\_\_\_\_\_ and was amended on

\_\_\_\_\_ (se applicabile)

\_\_\_\_\_ (if applicable)

Il sottoscritto dichiara inoltre di aver letto e compreso il contenuto della descrizione identificata in precedenza, rivendicazioni comprese, come modificati dall'eventuale modifica summenzionata

I hereby state that I have reviewed and understand the contents of the above identified specification, including the claims, as amended by any amendment referred to above

Il sottoscritto riconosce l'obbligo di rivelare informazioni essenziali ai fini della determinazione della brevettabilità ai sensi del Titolo 37, Codice dei Regolamenti Federali, § 1.56.

I acknowledge the duty to disclose information which is material to patentability as defined in Title 37, Code of Federal Regulations, § 1.56

## Italian Language Declaration

Il sottoscritto rivendica con la presente la priorità prevista dal Titolo 35, Codice degli Stati Uniti, § 119(e)(d) o § 365(b) in relazione a qualsiasi domanda o domanda estera di brevetto o certificato di inventore, o dal Titolo 35, § 365(a) degli stessi Codice in relazione a qualsiasi domanda internazionale PCT nella nazione in quale è designato almeno un paese diverso dagli Stati Uniti, i suddetti domande e certificati essendo elencati sotto, e, spuntando le seguenti domande, ha anche identificato sotto qualsiasi domanda estera di brevetto o certificato di inventore, o domanda internazionale PCT, la cui data di deposito preceda quella dalla domanda per la quale è rivendicata la priorità.

I hereby claim foreign priority under Title 35, United States Code, § 119(a)-(d) or § 365(b) of any foreign application(s) for patent or inventor's certificate, or § 365(a) of any PCT International application, which designated at least one country other than the United States, listed below, and have also identified below, by checking the box, any foreign application for patent or inventor's certificate, or PCT International application having a filing date before that of the application on which priority is claimed.

Prior Foreign Application(s)  
(Demande Estere Anteriori)

Priority claimed  
Diritto di priorità  
rivendicato

9823871.0 Great Britain  
(Number) (Country)  
(Número) (Nazione)

30 October 1998  
(Day/Month/Year Filed)  
(Giorno/Mese/Anno di deposito)

Yes  
 No

(Number) (Country)  
(Número) (Nazione)

(Day/Month/Year Filed)  
(Giorno/Mese/Anno di deposito)

Yes       No

Il sottoscritto rivendica con la presente i benefici previsti dal Titolo 35, Codici degli Stati Uniti, § 119(e), in relazione a qualsiasi domanda o domande provvisorie degli Stati Uniti elencate sotto.

I hereby claim the benefit under Title 35, United States Code, § 119(e) of any United States provisional application(s) listed below.

(Application No.)  
(Nº della domanda)

(Filing Date)  
Data di deposito)

(Application No.)  
(Nº della domanda)

(Filing Date)  
(Date d'admission)

Il sottoscritto rivendica con la presente i benefici previsti dal Titolo 35, Codice degli Stati Uniti, § 120, in relazione a qualsiasi domanda o domande statunitensi, o dal Titolo 35, § 365(c) degli stessi Codici in relazione a qualsiasi domanda internazionale PCT nella quale sono designati gli Stati Uniti, i suddette domande essendone elencate sotto e, nella misura in cui lo oggetto di ciascuna rivendicazione di questa domanda non sia stato espresso nella domanda statunitense o internazionale PCT anteriore, nel modo previsto dal primo paragrafo del Titolo 35, Codice degli Stati Uniti, § 112, riconosce l'obbligo di rivelare informazioni essenziali ai fini della determinazione della brevettabilità ai sensi del Titolo 37, Codici dei Regolamenti Federali, § 1.56, le quali diventano disponibili durante il periodo compreso tra la data di deposito della domanda anteriore e la data di deposito nazionale o internazionale PCT della presente domanda.

I hereby claim the benefit under Title 35, United States Code, § 120 of any United States application(s), or § 365(c) of any PCT-International application designating the United States, listed below and, insofar as the subject matter of each of the claims of this application is not disclosed in the prior United States PCT-International application in the manner provided by the first paragraph of Title 35, United States Code, § 112, I acknowledge the duty to disclose information which is material to patentability as defined in Title 37, Code of Federal Regulations, § 1.56 which became available between the filing date of the prior application and the earliest of PCT-International filing date of this application.

(Application No.)  
(Nº della domanda)

(Filing Date)  
(Data di deposito)

(Status) (patented, pending, abandoned) (Stato) (concessione di brevetto, in corso di esame, abbandono)

(Application No.)  
(Nº della domanda)

(Filing Date)  
(Data di deposito)

(Status) (patented, pending, abandoned)

Con la presente, il sottoscritto dichiara veritiera tutte le affermazioni contenute in questa domanda in relazione alle proprie conoscenze e di ritenere vere tutte le affermazioni o informazioni presentate. Le affermazioni o informazioni sono state espresse nella piena consapevolezza che le dichiarazioni enunciative false sono punibili con una pratica, l'incoraggiorne o entrarmi, ai sensi della Sezione 1001 del Titolo 18 del Codice degli Stati Uniti, e che tali dichiarazioni enunciative false possono mettere a repaginio la validità della domanda o di qualsiasi brevetto tascabile in merito.

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true, and further that these statements were made under the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.

## Italian Language Declaration

PROCURA: Il sootscritto inventore nomina con la presente il seguente avvocato o avvocati e/o agente o agenti al fine di istruire questa pratica e di condurre tutte le operazioni ad essa pertinente presso l'Ufficio dei Brevetti e Marchi di Fabbrica: (Elencare il nome ed il numero di matricola).

Norman F. Oblon, Reg. No. 24,618; Marvin J. Spivak, Reg. No. 24,913; C. Irvin McClelland, Reg. No. 21,124; Gregory J. Maier, Reg. No. 25,599; Arthur I. Neustadt, Reg. No. 24,854; Richard D. Kelly, Reg. No. 27,787; James D. Hamilton, Reg. No. 28,421; Eckhard H. Kuesters, Reg. No. 28,870; Robert T. Pous, Reg. No. 29,099; Charles L. Gholz, Reg. No. 26,395; William E. Beaumont, Reg. No. 30,986; Jean-Paul Lavallee, Reg. No. 31,451; Stephen G. Baxter, Reg. No. 32,884; Richard L. Treanor, Reg. No. 36,329; Steven P. Wehrhouch, Reg. No. 34,426; James J. Kulbaski, Reg. No. 34,648; Richard A. Neifeld, Reg. No. 35,299; J. Derek Mason, Reg. No. 35,270; Surinder Sachar, Reg. No. 34,423; Christina M. Gadiano, Reg. No. 37,628; Jeffrey B. McInlyre, Reg. No. 36,867; William T. Enos, Reg. No. 33,128; Michael E. McCabe, Jr., Reg. No. 37,182; Bradley D. Lytle, Reg. No. 40,073; and Michael R. Casey, Reg. No. 40,294, with full powers of substitution and revocation.

POWER OF ATTORNEY As a named inventor, I hereby appoint the following attorney(s) and/or agent(s) to prosecute this application and transact all business in the Patent and Trademark Office connected therewith: (list name and registration number)

Inviare le corrispondenza a:

Send Correspondence to  
OBLON, SPIVAK, McCLELLAND, MAIER & NEUSTADT, P.C.  
 FOURTH FLOOR  
 1755 JEFFERSON DAVIS HIGHWAY  
 ARLINGTON, VIRGINIA 22202 U.S.A.

Telefonare a:  
 (Nome e numero telefonico)

Direct Telephone calls to (name and telephone number)  
 (703) 413-3000

Nome e cognome dell'unico o del primo inventore

1/0 Name of sole or first inventor  
Paolo Peverello

Firma dell'inventore

Data

Inventor's signature  
Paolo Peverello

Date

12 March 2001

Residenza

Residence

Pavia, Italy ITX

Cittadinanza

Citizenship

Italian

Recapito postale

Post Office Address

Piazza San Pietro in Ciel d'Oro 7/A,  
27100 Pavia, Italy

Nome e cognome dell'eventuale secondo coinventore

2/0 Full name of second joint inventor, if any

Raffaella Amici

Firma del secondo coinventore

Data

Second inventor's signature

Date

Raffaella Amici 12 March 2001

Residenza

Residence

Piacenza, Italy ITX

Cittadinanza

Citizenship

Italian

Recapito postale

Post Office Address

Via N. Rocca 11, 29100 Piacenza, Italy

(Fornire le stesse informazioni e le firme del terzo e degli ulteriori coinventori)

(Supply similar information and signature for third and subsequent joint inventors)

### Italian Language Declaration

Nome per intero di un eventuale terzo co-inventore		Full name of third joint inventor, if any	
Firma del Terzo Inventore	Data	Third inventor's signature	Date
Residenza		Residence	Milan, Italy <i>ITX</i>
Cittadinanza		Citizenship	Italian
Recapito postale		Post Office Address	Via F. Cilea 106, 20151 Milan, Italy
Nome per intero di eventuale quarto co-inventore <i>4/50</i>		Full name of fourth joint inventor, if any	
Firma Quarto Inventore	Data	Fourth inventor's signature	Date
Residenza		Residence	Lurago d'Erba (Como), Italy <i>ITX</i>
Cittadinanza		Citizenship	Italian
Recapito postale		Post Office Address	Via San Bernardino 12, 22040 Lurago
Nome per intero di un eventuale quinto co-inventore <i>5/50</i>		Full name of fifth joint inventor, if any	
Firma Quinto Inventore	Data	Fifth inventor's signature	Date
Residenza		Residence	Brugherio (Milan), Italy <i>ITX</i>
Cittadinanza		Citizenship	Italian
Recapito postale		Post Office Address	Via Volturno Portici/2 80, 20047
Nome per intero di un eventuale sesto co-inventore <i>6/60</i>		Full name of sixth joint inventor, if any	
Firma del Sesto Inventore	Data	Sixth inventor's signature	Date
Residenza		Residence	Milan, Italy <i>ITX</i>
Cittadinanza		Citizenship	Italian
Recapito postale		Post Office Address	Via Montecatini 14, 20144 Milan, Italy

(Si prega di fornire simili informazioni e firme per il terzo e gli eventuali ulteriori co-inventori )

(Supply similar information and signature for third and subsequent joint inventors )